

(1 of 90)

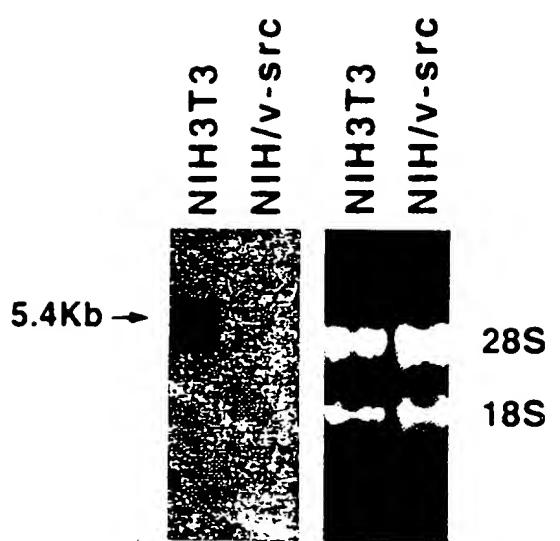


FIG. 1

(2 of 90)

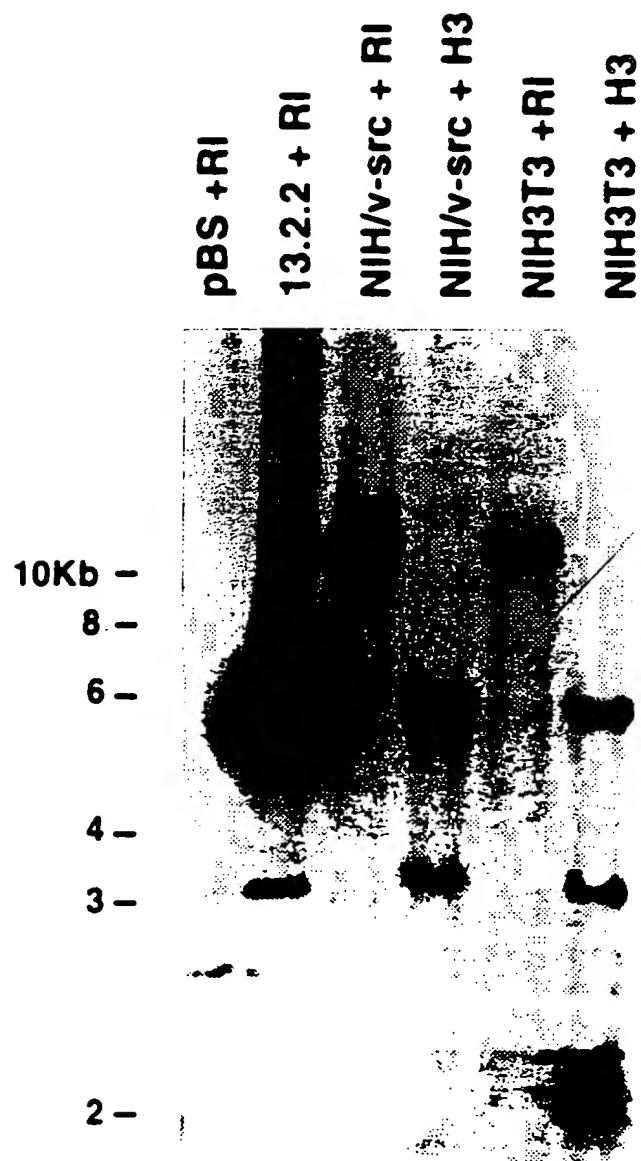
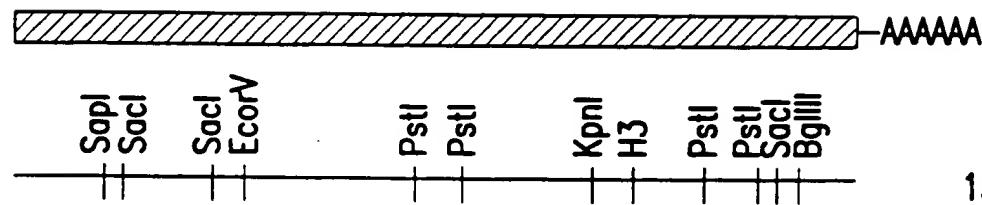


FIG.2A

(3 of 90)

322 cDNA (5.4Kb)



13.22

FIG.2B

(4 of 90)

	gaaaaagacagccaggaggccctcgaggaggcaggagccggcagaaggacacagaccaggccag	60
	gttgtcaggcagactacagaagggtggagctgcccattggaaaccaggltgtgacctgtga	120
	gycatcgtagggagaatgtgtgtccattttggcaacggaaagtgtttgtgaaatggatggat	180
	M E	2
181	agcccacccaaagaagtgttgcaagggtccacgtggacccgtggagaaggacagaggaggaa	240
3	A H Q F V V A E V S T V E K T E E E	22
241	qcaggaggaggaggaggctgaaagggggggtgggtggtagaaggaaacaggaaatccctt	300
23	Q <u>G</u> <u>G</u> E A E <u>G</u> V V V E G 1 G E S L	42
301	gccccctgaaagaaactgtgtggcccccaggagggtcccccaggaaaggctggcgtggat	360
43	P P E K I A E <u>P</u> Q F V P Q E A E P A E E	62
361	gctgtggaaaggcagaggatgtgtctctggaggagaccacactcaactgacagacact	420
63	L M K S R E M C V E G G D H T Q L T D L	82
421	aagtccctgaaaggacgtgtcccaaaacaccaggaaaggcattgtcaagtggat	480
83	S P E E R T L P K H P E G I V S E V E M	102
481	gctgtccctcaggaaagaatcaaggatcacaggaaagtcccttgaagaaactcttcgttag	540
103	L S S Q E R I K V Q G S P L K K L F S S	122
541	ctcaggcttaaggactgtctggaaagaaggcagaaggaaaacgaggagggtggggaga	600
123	S G L K K L S G <u>K</u> <u>Q</u> <u>K</u> G K R <u>G</u> <u>G</u> D	142
601	cgaagagccctggagaatccaacacattcacaccgaatccccagaggtgtgatgagaca	660
143	E E P G E Y Q H I H T E S P E S A D E Q	162

FIG.3A

(5 of 90)

661	gaaggaggaggagctctcgltcgccccggaggaccacgtgtctggagaa	720
163	K G E S A S P E E P E T T C L E K	182
721	aggccgcctggaaaggccccaggatggggaaagctggaaactacttcgtggagaa	780
183	G P L E A P R M G K L R K E L L R G E <u>K</u>	202
781	gaaggaggaggatcaactccctggcatccttcaaaaatgtgtgacaccacaaacgt	840
203	<u>K R K</u> D H S L G I L Q K D G D T Q E T V	222
841	ccgaaggaccttctgaggtgacaaggaggaaagactggagaaggctcaagagcgcaccc	900
223	R R P S F S D K F E L F K V K S A I L	242
901	glccctccacatgtatgcacatgtcagaaatlgcaagatgaagtcaaaaactgttgtgaggaa	960
243	S S T D S T V S E M Q D E V K T V G E E	262
961	acaaaaggccaggaaaccaaaggcgtaagggtgtgatacttcagtgcttgtggaaagcactgtat	1020
263	Q K P E P K R R V D T S V S W E A L I	282
1021	tttgtgtcgatcatccaaaggaggcaaggaggcatccatcttcagatataaaggaggcc	1080
283	C V G S S K <u>K R A R</u> K A S S D I R G P	302
1081	aggacacactggggggacagtacagaggcaggaggccaggcaaaagaaggcc	1140
303	R T L G G Q S O S R G G Q Q R S R	322
1141	aaacagacgtgttccgtccaggaccacggaggcaggaccacggcaaggaaagttcacc	1200
323	T D A V P A S T Q E Q D Q A Q G S S S P	342
1201	cggccaggggaaaggcccttcggaaagggtgtctccacttggagtcatttaaaag	1260
343	E P A G S P S E G V S T W E S F K R	362

FIG. 3B

(6 of 90)

1261	attagtcaactccaagaaaaaatccaagtcaaaaaacttggaaaggaaaaaggccggaaaggac	1320
363	L V T P R [K K S K] S K L E E K E A G R T	382
1321	tctagttgttagggagggttgtccacttagatcgAACCGTgttagagaatcttgggtt	1380
383	L V V G A G C P L R S N R V E K N L G F	402
1381	tccattaaagaat!catccccggacggggaaaaaggccatggggaaaggccaaagaaca	1440
403	P L R N S S P D G G R K G Q M G R Q E Q	422
1441	agccacttgtggaaaggacttcaggcccgagtggagataaatggggacggccgtatgtcccagc	1500
423	A T V E D S G P V E I N E D E P D V P A	442
1501	agtctgtccctctgtctgagttatgtcagttggagggaaaggatgtggaaaggccaggggaa	1560
443	V V P L S E Y D A V E R E K M E A Q G N	462
1561	tgcggaggctggcccgactgtctgggtgttagtgtccggaggctcagtaagaactctgtt	1620
463	A E L P S C W G C V V S E E L S K T L V	482
1621	ccacacttgtggaggcttcattgtggaccaggccgttccagggttcggaaaggcg	1680
483	H T V S V A V I D G T R A K T S K E E R	502
1681	gttctcccttggatatcccgcttccgttaacagaacacacacaggccggaaaggc	1740
503	S P S W I S A S V T E P L E H T A G E A	522
1741	catgcccacccgttgtggagggtcaactgtggaaaggacatccatgtggaaactccctgtgt	1800
523	M P P V E E V T E K D I I A E E T P V L	542
1801	cacccagaggttaccagggttaaaggatgccatgacgacatggtaaccaggtaatggaa	1860
543	T Q T L P E G K D A H D M V T S E V D	562

FIG. 3C

1861 563	tttacacctagaactgtgtacagccacagagacctcagaggcttcgtactgaagaagt F T S E A V T A T E T S E A L R T E E V	1920 582	
1921 583	tacccaaagcatcgggcccgaaagacccacagacatgggttcgcgtttcccaactgac T E A S G A E T T D M V S A V S Q L T	1980 602	
1981 603	tgactcccaagacaccacagggaaagccaccacccaggtaaggaggtagagggtgggtgt D S P D T T E A T p V Q E V E G G V L	2040 622	
2041 623	agatacayaagaagagqaycgcacccaggccatccctccaaaggccgttgcagacaagg D T E F F R Q T Q A T I I Q A V A D K V	2100 642	
2101 643	gaaaggagggtcccaagttccctgcacccaggactgtgcagagaacgggtcaaaagca K L T S Q V p A T Q T V Q R T G S K A L	2160 662	
2161 663	ggagaagggttggaggtagaggaggactccgaaggactggcttcggagaaaaaggaaagg F K V E E D S E V L A S E K E K D	2220 682	
2221 683	cgttatgccaaaggaccggctgtggaggactggatctggatcttgcacagggtctga V M P K G P V Q E A G A E H L A Q G S E	2280 702	
2281 703	gactggacaggctactccaggaggcttgaagttccgtacagcagatgtagacca T G Q A T P E S L E V P E V T A D V D H	2340 722	
2341 723	tgtcgccacgtgccaggttatcaaggctccaggactgtggaaacaggccgtggccctgt V A T C Q V I K L Q Q L M E Q A V A P E	2400 742	
2401 743	gtcatccgaaacctttagacagacactgtgagacaatggaaacggactccctttagcattc S S E T L T D S E T N G S T R L A D S D	2460 762	

(7 of 90)

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FIG. 3D

(8 of 90)

2461 763	cactgcagatggcacacaaggatgaaaccattgcacggacacttgcaggactaaaggccactgc T A D G T Q Q D E T I D S Q D S K A T A	2520 782
2521 783	agctgtcaggcagtcacaggtcacagaagaaggcggtactgtctcagaaaaggaggagcc A V E Q S Q V T E E A A T A Q K E E P	2580 802
2581 803	ttcgacactaataatgttccagccaggaaacaatggaaagaaccaggaaaggaga S T L P N N V P A Q E E H G E P G R D	2640 822
2641 823	tgttcttgaatctacacacgcaaggatgtctgtcaggccgtgcccgtctggcaaaagac V L E P T Q Q E L A A A V P V W Q K T	2700 842
2701 843	ttaggtggtcaaggagggtgactggatggatggatggaaaaggtaaaggaaaca E V G Q E G L V D W L D G E K V E E Q	2760 862
2761 863	ggagggttgttacactctggacccaaacagtcaaaaggctgtctgtacatgtacag E V F V H S G P N S Q K A A D V T Y D S	2820 882
2821 883	tgaagtgtatggatgtggccgggtcaggaaaaggaggtactgtcaggatcttag E V M G V A G C Q E K E S T E V Q S L S	2880 902
2882 903	cctggaggaggatggaaactgtacgttggaaaaggagacaaggccaga L E E G M E T D V E K E K R E T K P E	2940 922
2941 923	gcaagtgtatggatggcaggaaacaggatgtggaaaaggagactacgg Q V S E E G E Q E T A A P E H E R N Y G	3000 942
3001 943	gaaggccaggatgtacatgtggatgtggggggaaaggcactggggaaaggcc K P V L T L D M P S S E R G K A L G S L	3060 962

FIG.3E

(9 of 90)

3061 963	ttggagaaaggcccttctccaggaccaagacaagcaggttgcataaggltcaagttca G G S P S I P D Q D K A G C I E V Q V Q	3120 982
3121 983	aaggcctggacacaaacagtcactaaaaacagcagaaggctgtggaaaaaggatcatagaaaacgg S L D T V T Q T A E A V E K V I E T V	3180 1002
3181 1003	tgtgatttcagagacagggtgaaaatgtccaggatgtgttaggtgcacacattaccaggctga V I S E T G E S P E C V G A H L L P A E	3240 1002
3241 1023	ggaaatccttgtaaacgggtggccactggactctcagatgcagaggacacggatcccc K S S A T G G H W T L Q H A E D T V P L	3300 1042
3301 1043	ggggcctgtggatctcaggcagaatccatccatcatgttaactccgtccctgtggaaaggcac G P F S Q A F S I P I I V T P A P E S T	3360 1062
3361 1063	cctacatccgtacaaaggaaataaggcatcccaaggatcgatcgaggaaaga I H P D L Q G E I S A S Q R E R S E E	3420 1082
3421 1083	ggacaaggccagatgtggcgtatgtggcaaggaaaggatcagcaatcgacaaatgt D K P D A G P D A D G K E S T A I D K V	3480 1102
3481 1103	cctcaaggctgaaacctgtggaaacttggatgtggatggaaaggaaatgtgtgt L K A E P E I L E L E S K S N K I V L N	3540 1122
3541 1123	cgcattcagacaggccgtgtaccaggttcgcaatcgatcagaaaacaggcccccaactcatgc V I Q T A V D Q F A R T E I A P E T H A	3600 1142
3601 1143	ttatgattcacagaccagggttcctgcaatgcgcgttggacaggagccaaacagatg Y D S Q T Q V P A M R L D S R E P N R C	3660 1162

FIG.3F

3661	ctggacaaaaatgaaatggccaaagatgaaacacccagt.gccgcaggccaggaggactt	3720
1163	W T K M V A K M K H P V P R E D L	1182
3721	gcaaggcctgaccgttctggggcatggctcagtcgaaaatgtcttgcgcgttgca	3780
1183	Q V L T V L E A W L S S E M L A A L A V	1202
3781	tggaaagcgcgggtcaaaagttaaggcattgagaagctgtcctcaacccaaaggataaaa	3840
1203	F S A G V K V S I E K L P P Q P K D Q K	1222
3841	ggagcatgtgtgtatggccctcaagtcgtccaaaggcttaaggccaggcaggatgtclgg	3900
1223	F H A D G P Q L Q S I A Q A E V S G	1242
3901	aaacctaacaaaaatccccagacaccacccaaacggaccataacggaggatgtcccc	3960
1243	N I I K F S P D I N G P K L T E E R C P	1262
3961	ccaaaaatgtgggtccaggaaaggaaatgtctaccaaaggatgtcgttcaaaaggaaaggcc	4020
1263	Q K L R S R K K C L P S Q R T R P	1282
4021	caggcagaaggaggaccgtgcaggaggccaaaggagacctggcagaatcctaaggatgttagt	4080
1283	R Q K R T C R S Q R E T W Q N P K M L V	1302
4081	tgctcattqtacatgttaagaccagaatgtaaaaacaaggatcacagaaacaaggatgtctgt	4140
1303	A H C T S V R P E C E N K S Q N K M L L	1322
4141	gttgggaccattggaccaaggatttcagagccatgagatccaggagagcaggccgtccaat	4200
1323	L G P W T K I S E P M R S R E Q G R P M	1342
4201	gatttccacccaggtagagcacccgacaattctggggctcatcggagactagagccagc	4260
1343	I S T Q *	1346

(10 20)

90

(11 of 90)

FIG. 3H

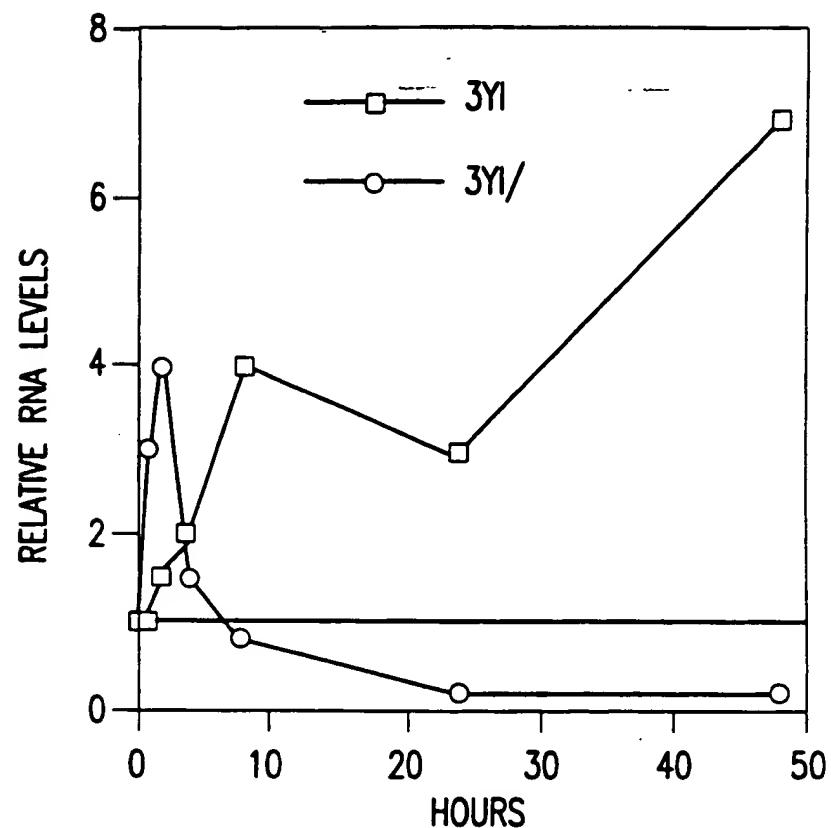


FIG.4A

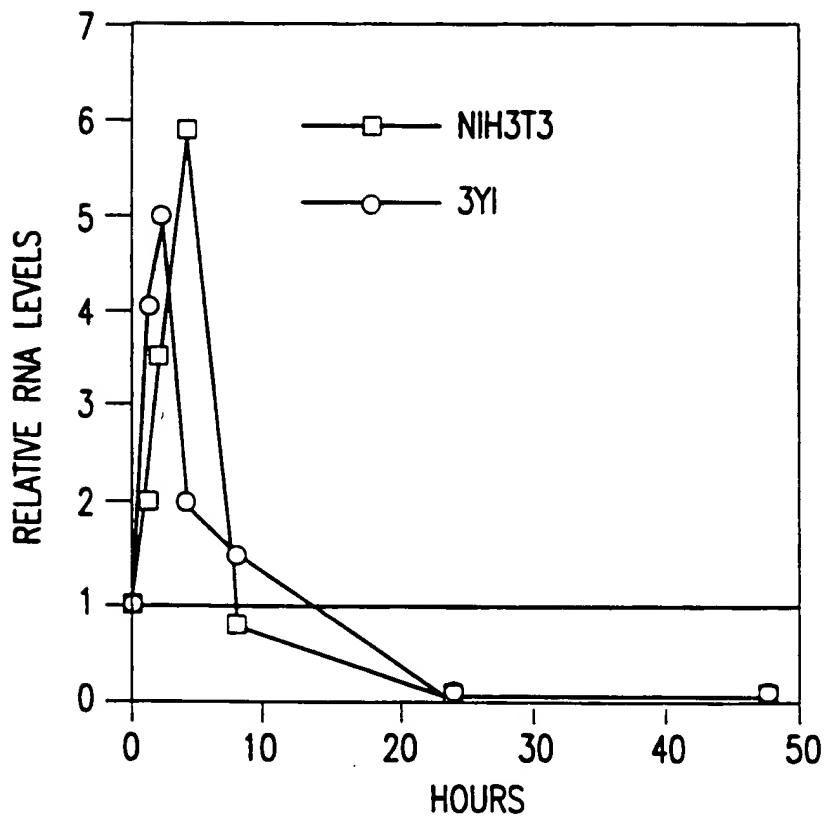


FIG.4B

(13 of 90)

rat-6
rat-6/raf-1
rat-6/raf5
rat-6/neu
rat-6/myc
rat-6/src
rat-6/mos



FIG.5

(14 of 90)

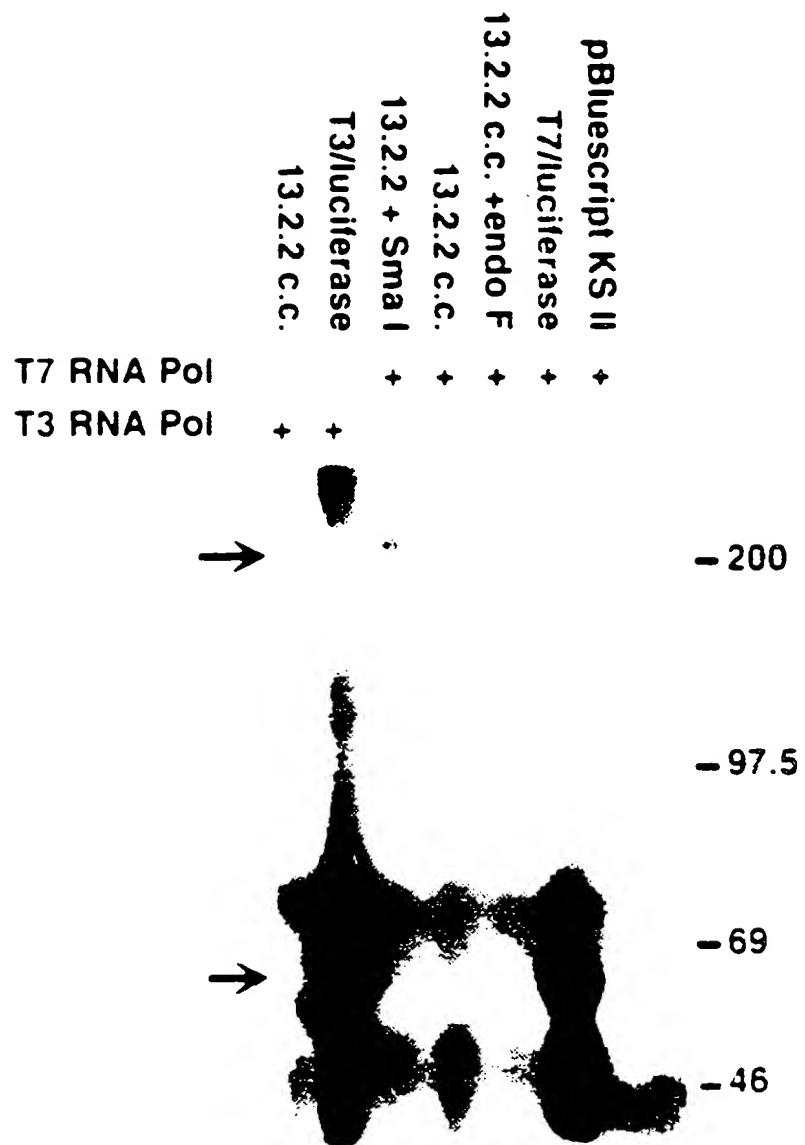


FIG.6

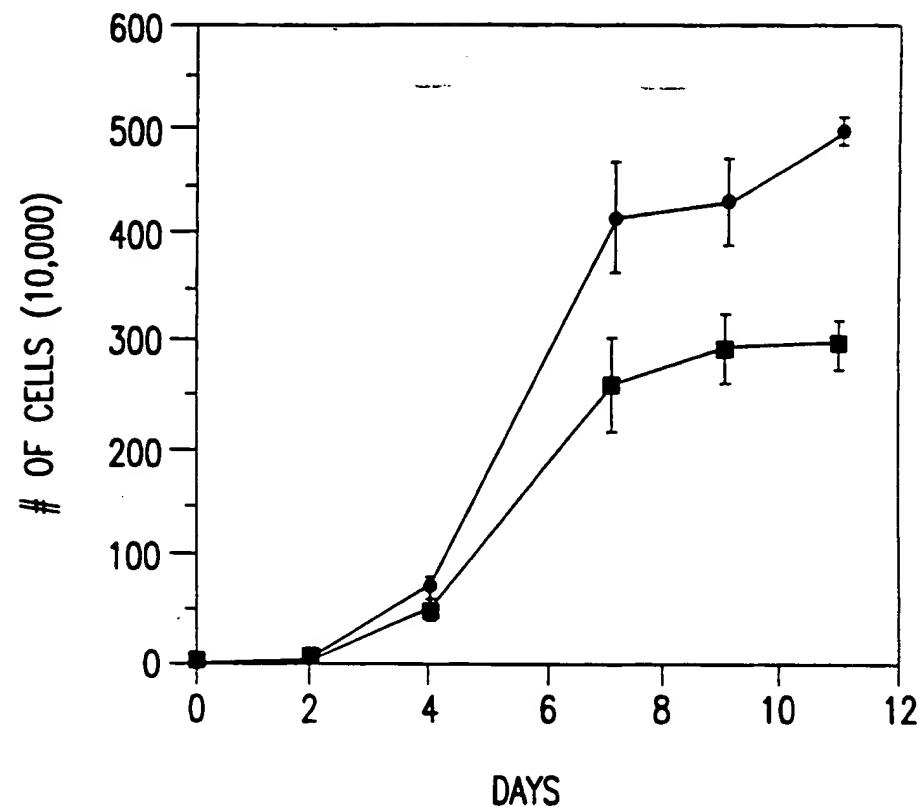


FIG.7A

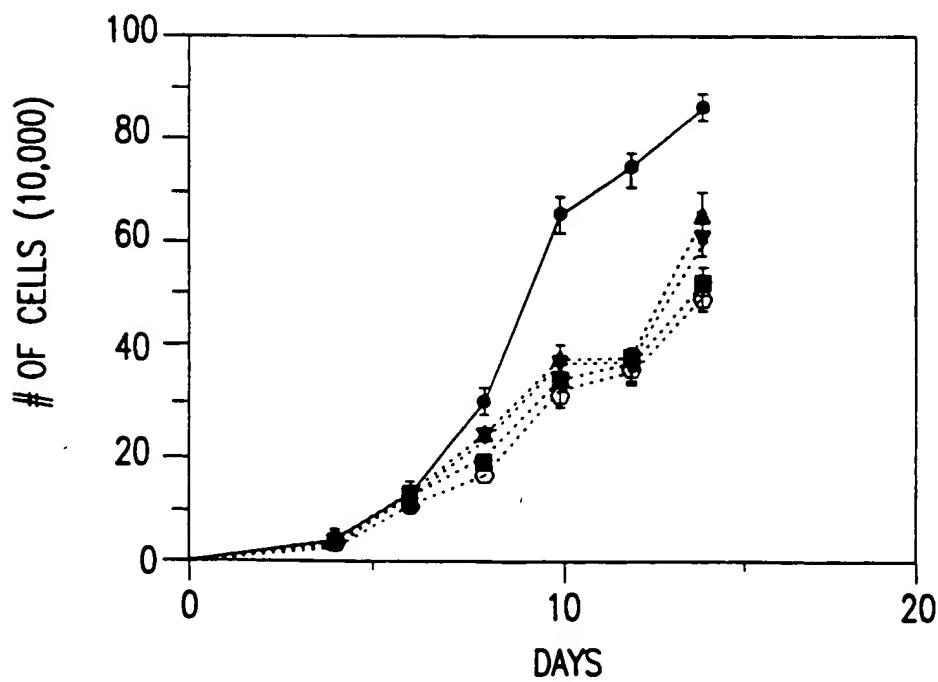


FIG.7B

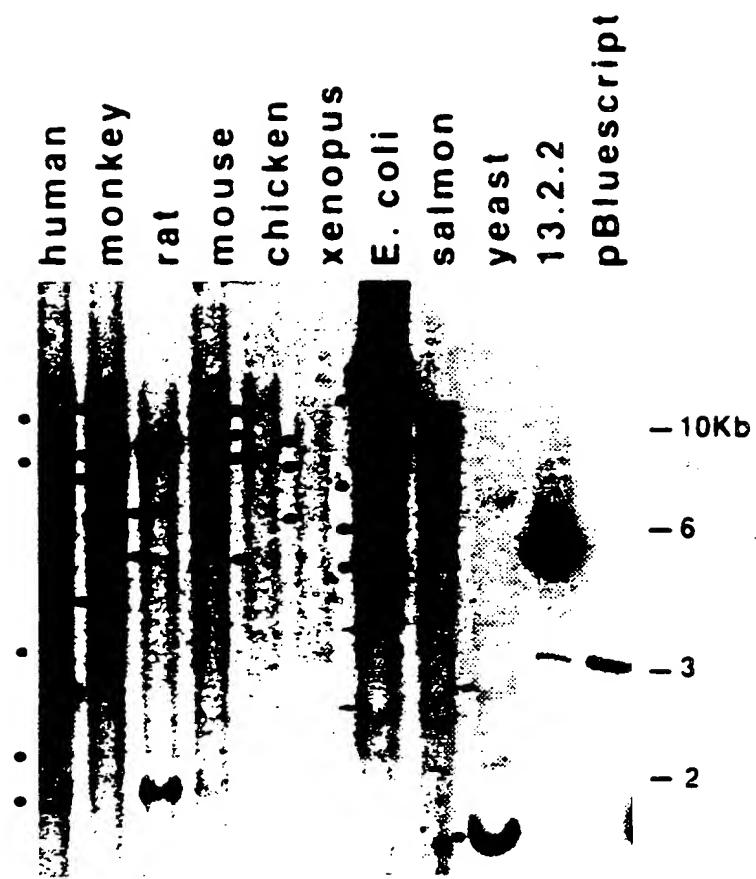


FIG. 8

(17 of 90)

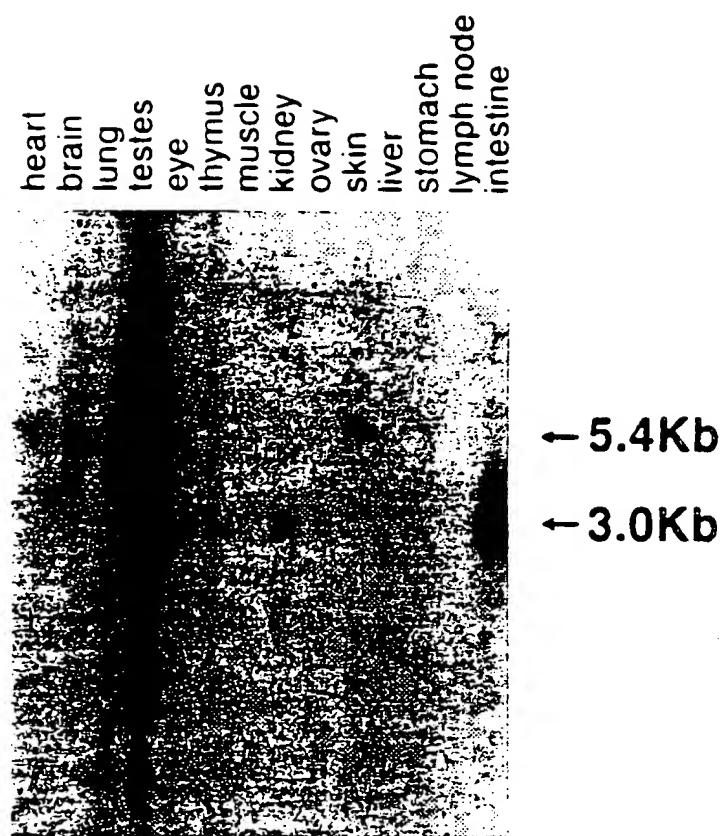


FIG.9

(19 of 90)

9 18 27 36 45 54

5' ATG GGC GCA GGC AGT TCC ACC GAG CAG CGG AGC CCC GAG CAG CCG GCG GGG AGC

M G A G S S T E Q R S P E Q P A G S

63 72 81 90 99 100

GAC ACG CCG AGC GAG CTG GTG CTC AGT GGC CAT GGG CCC GCA GCT GAA GCC TCG

D T P S E L V L S G H G P A A E A S

117 126 135 144 153 162

GGA GCA GCT GGA GAC CCC GCC GAC GCG GAC CCC GCC ACC AAG CTC CCA CAG AAG

G A A G D P A D A D P A T K L P Q K

171 180 189 198 207 216

AAT GGC CAG CTG TCT TCT GTC AAC GGC GTA GCT GAA CAA GGA GAT GTC CAT GTC

N G Q L S S V N G V A E Q G D V H V

225 234 243 252 261 270

CAA GAG GAA AAC CAG GAG GGG CAG GAG GAA GAA GTC GTT GAT GAG GAT GTT GGA

Q E E N Q E G Q E E E V V D E D V G

279 288 297 306 315 324

CAG CGA GAG TCA GAA GAT GTG AGA GAA AAA GAC CGA GTT GAA GAA ATG GCG GCC

Q R E S E D V R E K D R V E E M A A

333 342 351 360 369 378

AAC TCC ACA GCT GTT GAA GAT ATC ACA AAG GAT GGG CAG GAG GAG ACA TCA GAA

N S T A V E D I T K D G Q E E T S E

387 396 405 414 423 432

ATA ATT GAA CAG ATC CCT GCT TCA GAA AAC AAT GTG GAA GAA ATG GTA CAG CCT

I I E Q I P A S E N N V E E M V Q P

FIG.11A

(20 of 90)

441 450 459 468 477 486
GCT GAG TCC CAG GCT AAT GAT GTT GGC TTC AAG AAA GTA TTT AAA TTT GTT GGT

A E S Q A N D V G F K K V F K F V G

495 504 513 522 531 540
TTT AAA TTC ACG GTG AAG AAG GAT AAA AAT GAA AAG TCA GAT ACT GTC CAA CTA

F K F T V K K D K N E K S D T V Q L

549 558 567 576 585 594
CTC ACT GTC AAG AAG GAT GAA GGC GAA GGG GCA GAA GCC TCT GTC GGA GCT GGA

L T V K K D E G E G A E A S V G A G

603 612 621 630 639 648
GAC CAC CAG GAG CCC AGT GTG GAG ACT GCC GTC GGA GAG TCA GCA TCC AAA GAA

D H Q E P S V E T A V G E S A S K E

657 666 675 684 693 702
AGT GAG CTG AAG CAA TCC ACA GAG AAG CAA GAA GGC ACC CTG AAG CAA GAA CAG

S E L K Q S T E K Q E G T L K Q E Q

711 720 729 738 747 756
AGC AGC ACA GAA ATC CCC CTT CAA GCC GAA TCT GAT CAA GCG GCT GAG GAA GAA

S S T E I P L Q A E S D Q A A E E E

765 774 783 792 801 810
GCC AAA GAT GAA GGA GAA AAA CAA GAG AAA GAG CCC ACC AAG TCC CCA GAA

A K D E G E E K Q E K E P T K S P E

819 828 837 846 855 864
TCC CCG AGC AGC CCA GTC AAC AGT GAG ACA ACA TCT TCC TTC AAG AAG TTC TTC

S P S S P V N S E T T S S F K K F F

FIG.11B

(21 of 90)

873	882	891	900	909	918												
ACT	CAC	GGT	TGG	GCC	GGC	TGG	CGC	AAG	AAG	ACC	AGC	TTC	AAG	AAA	TCA	AAA	GAG
T	H	G	W	A	G	W	R	K	K	T	S	F	K	K	S	K	E
927	936	945	954	963	972												
GAT	GAT	CTG	GAA	ACT	GCC	GAG	AAG	AGA	AAG	GAG	CAA	GAG	GCA	GAA	AAA	GTA	GAC
D	D	L	E	T	A	E	K	R	K	E	Q	E	A	E	K	V	D
981	990	999	1008	1017	1026												
GAG	GAA	GAA	AAG	GAA	AAG	ACA	GAG	CCA	GCC	TCG	GAG	GAG	CAG	GAG	CCG	GCA	GAA
E	E	E	K	E	K	T	E	P	A	S	E	E	Q	E	P	A	E
1035	1044	1053	1062	1071	1080												
GAC	ACA	GAC	CAG	GCC	AGG	TTG	TCA	GCA	GAC	TAC	GAG	AAG	GTG	GAG	CTG	CCT	TTG
D	T	D	Q	A	R	L	S	A	D	Y	E	K	V	E	L	P	L
1089	1098	1107	1116	1125	1134												
GAA	GAC	CAG	GTT	GGT	GAC	CTG	GAG	GCA	TCG	TCA	GAG	GAG	AAG	TGT	GCT	CCT	TTG
E	D	Q	V	G	D	L	E	A	S	S	E	E	K	C	A	P	L
1143	1152	1161	1170	1179	1188												
GCA	ACG	GAA	GTG	TTT	GAT	GAG	AAG	ATG	GAA	GCC	CAC	CAA	GAA	GTT	GTT	GCA	GAG
A	T	E	V	F	D	E	K	M	E	A	H	Q	E	V	V	A	E
1197	1206	1215	1224	1233	1242												
GTC	CAC	GTG	AGC	ACC	GTG	GAG	AAG	ACA	GAG	GAG	GAG	CAG	GGA	GGA	GGA	GAG	
V	H	V	S	T	V	E	K	T	E	E	E	Q	G	G	G	G	E
1251	1260	1269	1278	1287	1296												
GCT	GAA	GGG	GGC	GTG	GTG	GTA	GAA	GGA	ACA	GGA	GAA	TCC	TTG	CCC	CCT	GAG	AAA
A	E	G	G	V	V	V	E	G	T	G	E	S	L	P	P	E	K

FIG.11C

(22 of 90)

1305 1314 1323 .. 1332 1341 1350
CTG GCT GAG CCC CAG GAG GTC CCC CAG GAA GCT GAG CCT GCT GAG GAG CTG ATG

L A E P Q E V P Q E A E P A E E L M

1359 1368 1377 1386 1395 1404
AAG AGC AGA GAG ATG TGT GTC TCT GGA GGA GAC CAC ACT CAA CTG ACA GAC CTA

K S R E M C V S G G D H T Q L T D L

1413 1422 1431 1440 1449 1458
AGT CCT GAA GAG AAG ACG CTG CCC AAA CAC CCA GAA GGC ATT GTC AGT GAG GTG

S P E E K T L P K H P E G I V S E V

1467 1476 1485 1494 1503 1512
GAG ATG CTG TCC TCT CAG GAA AGA ATC AAG GTA CAG GGA AGT CCC TTG AAG AAA

E M L S S Q E R I K V Q G S P L K K

1521 1530 1539 1548 1557 1566
CTC TTC AGT AGC TCA GGC TTA AAG AAG CTG TCT GGG AAG AAG CAG AAG GGG AAA

L F S S S G L K K L S G K K Q K G K

1575 1584 1593 1602 1611 1620
CGA GGA GGT GGG GGA GAC GAA GAG CCT GGA GAA TAC CAA CAC ATT CAC ACC GAA

R G G G G D E E P G E Y Q H I H T E

1629 1638 1647 1656 1665 1674
TCC CCA GAG AGT GCT GAT GAG CAG AAG GGA GAG AGC TCT GCG TCG TCC CCC GAG

S P E S A D E Q K G E S S A S S P E

1683 1692 1701 1710 1719 1728
GAG CCT GAG GAG ACC ACG TGT CTG GAG AAA GGG CCG CTG GAA GCA CCC CAG GAT

E P E E T T C L E K G P L E A P Q D

FIG.11D

1737	1746	1755	1764	1773	1782
GGG GAA GCT GAG GAA GGA ACT ACT_TCC GAT GGA GAG_AAG AAG AGA GAA GGG ATC					
G E A E E G T T S D G E K K R E G I					
1791	1800	1809	1818	1827	1836
ACT CCC TGG GCA TCC TTC AAA AAG ATG GTG ACA CCC AAG AAA CGG GTC CGA AGA					
T P W A S F K K M V T P K K R V R R					
1845	1854	1863	1872	1881	1890
CCT TCT GAG AGT GAC AAG GAG GAA GAG CTG GAG AAG GTC AAG AGC GCC ACC TTG					
P S F S D K E E E L E K V K S A T L					
1899	1908	1917	1926	1935	1944
TCC TCC ACT GAT AGC ACA GTG TCA GAA ATG CAA GAT GAA GTC AAA ACT GTT GGT					
S S T D S T V S E M Q D E V K T V G					
1953	1962	1971	1980	1589	1998
GAG GAA CAA AAG CCA GAG GAA CCA AAG CGT AGG GTG GAT ACT TCA GTG TCT TGG					
E E Q K P E E P K R R V D T S V S W					
2007	2016	2025	2034	2043	2052
GAA GCA CTG ATT TGT GTC GGA TCA TCC AAG AAG AGA GCA AGG AAG GCA TCC TCT					
E A L I C V G S S K K R A R K A S S					
2061	2070	2079	2088	2097	2106
TCA GAT GAT GAA GGA GGG CCA AGG ACA CTG GGA GGG GAC AGT CAC AGA GCA GAG					
S D D E G G P R T L G G D S H R A E					
2115	2124	2133	2142	2151	2160
GAG GCC AGC AAA GAC AAA GAA GCC GGA ACA GAC GCT GTT CCT GCC AGC ACC CAG					
E A S K D K E A G T D A V P A S T Q					

FIG.11E

(24 of 90)

2169 2178 2187 2196 2205 2114
GAG CAG GAC CAA GCG CAA GGA AGT ICC TCA CCC GAG CCA GCG GGA AGC CTT TCC

E Q D Q A Q G S S S P E P A G S P S

2223 2232 2241 2250 2259 2268
GAA GGG GAA GGT GTC TCC ACT TGG GAG TCA TTT AAA AGA TTA GTC ACT CCA AGA

E G E G V S T W E S F K R L V T P R

2277 2286 2295 2304 2313 2322
AAA AAA TCC AAG TCA AAA CTG GAA GAG AAA GCC GAA GAC TCT AGT GTA GAG CAG

K K S K S K L E E K A E D S S V E Q

2331 2340 2349 2358 2367 2376
TTG TCC ACT GAG ATC GAA CCG AGT AGA GAA GAA TCT TGG GTT TCC ATT AAG AAA

L S T E I E P S R E E S W V S I K K

2385 2394 2403 2412 2421 2430
TTC ATC CCC GGA CGG CGG AAG AAA AGG GCA GAC GGG AAG CAA GAA CAA GCC ACT

F I P G R R K K R A D G K Q E Q A T

2439 2448 2457 2466 2475 2484
GTG GAA GAC TCA GGG CCA GTG GAG ATA AAT GAG GAC GAC CCT AAT GTC CCA GCC

V E D S G P V E I N E D D P N V P A

2493 2502 2511 2520 2529 2538
GTC GTG CCT CTG TCT GAG TAT AAT GCA GTG GAG AGG GAG AAG ATG GAA GCC CAG

V V P L S E Y N A V E R E K M E A Q

2547 2556 2565 2574 2583 2592
GGG AAT ACG GAG CTG CCC CAG CTG CTG GGG GCT GTG TAC GTG TCC GAG GAG CTC

G N T E L P Q L L G A V Y V S E E L

FIG.11F

(25 of 90)

2601	2610	2619	2628	2637	2646
AGT AAG ACT CTG GTC CAC ACT GTG AGT GTC GCA GTC ATT GAT GGG ACC AGG GCA					

S	K	T	L	V	H
T	V	H	T	V	S
V	A	V	I	I	D
A	G	T	R	R	A

2655	2664	2673	2682	2691	2700
GTC ACC AGT GTC GAA GAG CGG TCT CCT TCG TGG ATA TCC GCT TCC GTA ACA GAA					

V	T	S	V	E	E
R	S	P	S	W	I
S	A	S	A	S	V
V	T	E	T	E	E

2790	2718	2727	2736	2745	2754
CCT CTT GAA CAC ACA GCG GGA GAA GCC ATG CCA CCT GTT GAA GAG GTC ACT GAA					

P	L	E	H	T	A
G	E	A	M	P	P
M	P	V	P	V	E
P	E	V	T	E	E

2763	2772	2781	2790	2799	2808
AAA GAC ATC ATT GCA GAA GAA ACT CCT GTG CTC ACC CAG ACG TTA CCA GAG GGT					

K	D	I	I	A	E
E	T	P	V	L	T
T	Q	Q	T	L	P
E	T	P	E	P	E
G	E	E	G	G	G

2817	2826	2835	2844	2853	2862
AAA GAT GCC CAT GAC GAC ATG GTC ACC AGT GAA GTG GAT TTC ACC TCA GAA GCT					

K	D	A	H	D	D
M	V	T	S	E	V
V	D	D	S	E	E
E	F	F	E	T	S
G	T	T	V	E	E

2871	2880	2889	2898	2907	2916
GTC ACA GCC ACA GAG ACC TCA GAG GCT CTC CGT ACT GAA GAA GTT ACC GAA GCA					

V	T	A	T	E	T
S	E	S	E	A	L
E	A	L	R	T	R
A	T	R	E	E	E
T	E	E	V	V	E
E	A	L	T	E	A

2925	2934	2943	2952	2961	2970
TCG GGG GCC GAA GAG ACC ACA GAC ATG GTG TCC GCA GTT TCC CAG CTG ACT GAC					

S	G	A	E	E	T
T	T	D	M	V	S
V	S	A	V	S	Q
S	Q	V	S	L	T
V	L	T	D	T	D

2979	2988	2997	3006	3015	3024
TCC CCA GAC ACC ACA GAG GAA GCC ACC CCA GTT CAG GAG GTA GAG AGT GGT GTG					

S	P	D	T	T	E
E	E	E	E	E	E
A	A	A	A	A	A
T	T	T	T	T	T
P	P	P	P	P	P
V	V	V	V	V	V
Q	Q	Q	Q	Q	Q
E	E	E	E	E	E
V	V	V	V	V	V
E	E	E	E	E	E
S	S	S	S	S	S
G	G	G	G	G	G
V	V	V	V	V	V

FIG.11G

3033	3042	3051	3060	3069	3078
CTA GAT ACA GAA GAA GAG GAG CGC CAG ACG CAG GCC ATC CTC CAA GCC GTT GCA					
-----	-----	-----	-----	-----	-----
L	D	T	E	E	E
				R	Q
				T	Q
				A	I
				L	Q
				A	V
					A
3087	3096	3105	3114	3123	3132
GAC AAG GTG AAA GAG GAG TCC CAG GTG CCT GCA ACC CAG ACT GTG CAG AGA ACG					
-----	-----	-----	-----	-----	-----
D	K	V	K	E	E
				S	Q
				V	P
				A	T
				T	Q
				V	Q
				R	T
3141	3150	3159	3168	3177	3186
GGG TCA AAA GCA CTG GAG AAG GTT GAG GAG GTA GAG GAG GAC TCC GAA GTG CTG					
-----	-----	-----	-----	-----	-----
G	S	K	A	L	E
				K	V
				E	E
				V	E
				E	D
				S	S
				E	V
				V	L
3195	3204	3213	3222	3231	3240
GCT TCG GAG AAA GAG AAG GAC GTT ATG CCG AAA GGA CCC GTG CAG GAA GCT GGA					
-----	-----	-----	-----	-----	-----
A	S	E	K	E	K
				D	V
				M	M
				P	P
				K	G
				G	P
				P	V
				V	Q
				E	A
				G	G
3195	3258	3267	3276	3285	3294
GCT GAG CAT CTT GCA CAG GGC TCT GAG ACT GGA CAG GCT ACT CCA GAG AGC CTT					
-----	-----	-----	-----	-----	-----
A	E	H	L	A	Q
				G	G
				S	E
				E	T
				T	G
				G	Q
				A	A
				T	T
				P	P
				E	E
				S	S
				L	L
3303	3312	3321	3330	3339	3348
GAA GTT CCT GAA GTC ACG GCA GAT GTA GAC CAT GTC GCC ACG TGC CAG GTT ATC					
-----	-----	-----	-----	-----	-----
E	V	P	E	V	T
				A	A
				D	D
				V	H
				D	V
				H	A
				V	T
				A	C
				T	Q
				C	V
				Q	I
3357	3366	3375	3384	3393	3402
AAG CTC CAG CAG CTG ATG GAA CAG GCC GTG GCC CCT GAG TCA TCC GAA ACC TTG					
-----	-----	-----	-----	-----	-----
K	L	Q	Q	L	M
				M	E
				E	Q
				Q	A
				A	V
				V	A
				A	P
				P	E
				E	S
				S	S
				S	E
				E	T
				T	L
3411	3420	3429	3438	3447	3456
ACA GAC AGT GAG ACA AAT GGA AGC ACT CCC TTA GCA GAT TCA GAC ACT GCA GAT					
-----	-----	-----	-----	-----	-----
T	D	S	E	T	N
					G
					S
					T
					P
					L
					A
					D
					S
					D
					T
					A
					D

FIG.11H

3465	3474	3483	3492	3501	3510
GGG ACA CAG CAA GAT GAA ACC ATT GAC	AGC CAG GAC AGT AAA	GCC ACT GCA GCT			
G T Q Q D E T I D S Q D S K A T A A					
3519	3528	3537	3546	3555	3564
GTC AGG CAG TCA CAG GTC ACA GAA GAA	GAG GCG GCT ACT GCT CAG AAA GAG GAG				
V R Q S Q V T E E E A A T A Q K E E					
3573	3582	3591	3600	3609	3618
CCT TCG ACA CTA CCT AAT AAT GTT CCA GCC CAG GAA GAA	CAT GGG GAA GAA CCA				
P S T L P N N V P A Q E E E H G E E P					
3627	3636	3645	3654	3663	3672
GGA AGA GAT GTT CTT GAA CCT ACA CAG CAA GAG CTT ACT GCT	GCA GCC GTG CCC				
G R D V L E P T Q Q E L T A A A A V P					
3681	3690	3699	3708	3717	3726
GTT CTG GCA AAG ACT GAG GTG GGT CAA GAG GGT GAG GTT GAC TGG TTG GAT GGA					
V L A K T E V G Q E G E V D W L D G					
3735	3744	3753	3762	3771	3780
GAA AAA GTC AAA GAA GAA CAG GAG GTG TTT GTA CAC TCT GGA CCC AAC AGT CAA					
E K V K E E Q E V F V H S G P N S Q					
3789	3798	3807	3816	3825	3834
AAG GCT GCT GAT GTG ACA TAT GAC AGT GAA GTG ATG GGA GTG GCC GGG TGT CAG					
K A A D V T Y D S E V M G V A G C Q					
3843	3852	3861	3870	3879	3888
GAA AAG GAG AGT ACT GAA GTG CAG AGT CTT AGC CTG GAG GAG GGA GAG ATG GAA					
E K E S T E V Q S L S L E E G E M E					

3897	3906	3915	3924	3933	3942
ACT GAC GTT GAA AAG GAG AAA AGG GAG ACA AAG CCA GAG CAA GTG AGT GAA GAA					

T	D	V	E	K	E
K	R	E	T	K	P
P	E	Q	V	S	E

3951	3960	3969	3978	3987	3996
GGT GAG CAG GAA ACA GCC GCT CCT GAG CAT GAA GGA ACC TAC GGG AAG CCA GTC					

G	E	Q	E	T	A
A	P	E	H	E	G
P	E	H	G	T	Y
K	P	V	G	K	P

4005	4014	4023	4032	4041	4050
CTG ACA CTT GAC ATG CCC AGC TCA GAG AGG GGG AAG GCA CTG GGA AGC CTT GGA					

L	T	L	D	M	P
M	P	S	S	S	E
P	S	E	R	G	K
S	L	A	L	L	G

4059	4068	4077	4086	4095	4104
GGA AGC CCT TCT CTC CCA GAC CAA GAC AAA GCA GGT TGC ATA GAG GTT CAA GTT					

G	S	P	S	L	P
S	P	D	Q	D	Q
P	S	K	A	K	G
S	L	A	G	A	C

4113	4122	4131	4140	4149	4158
CAA AGC CTG GAC ACA ACA GTC ACT CAA ACA GCA GAA GCT GTG GAA AAG GTC ATA					

Q	S	L	D	T	T
S	L	D	T	V	T
L	D	T	V	T	Q
D	T	V	V	T	T

4167	4176	4185	4194	4203	4212
GAA ACG GTT GTG ATT TCA GAG ACA GGT GAA AGT CCA GAG TGT GTA GGT GAC CAC					

E	T	V	V	I	S
T	V	V	I	S	E
V	V	I	S	E	T
V	I	S	E	T	G

4221	4230	4239	4248	4257	4266
TTA TTA CCA GCT GAG AAG TCC TCT GCA ACG GGT GGC CAC TGG ACT CTT CAG CAT					

L	L	P	A	E	K
L	L	P	A	E	K
P	A	E	K	S	S
A	E	K	S	S	A

4275	4284	4293	4902	4311	4320
GCA GAG GAC ACG GTA CCC CTG GGG CCT GAG TCT CAG GCA GAA TCC ATC CCA ATC					

A	E	D	T	V	P
E	D	T	V	P	L
D	T	V	P	L	G
T	V	P	L	G	P
V	P	L	G	P	E
P	L	G	P	E	S
L	G	P	E	S	Q
G	P	E	S	Q	A
P	E	S	Q	A	E
E	S	Q	A	E	S
S	Q	A	E	S	I
Q	A	E	S	I	P
A	E	S	I	P	I

4329	4338	4347	4356	4365	4374
ATA	GTA	ACT	CCT	GCT	CCT
GAA	AGC	ACC	CTA	CAT	CCT
GAC	CTA	CAA	GGA	GAA	ATA
I	V	T	P	A	P
E	S	T	L	H	P
D	L	Q	G	E	I
4383	4392	4401	4410	4419	4428
AGC	GCA	TCC	CAG	AGA	GAG
CGA	TCA	GAG	GAA	GAG	GAC
AAG	CCA	GAT	GCT	GGT	CCT
S	A	S	Q	R	E
R	S	E	E	E	D
K	P	D	A	G	P
4437	4446	4455	4464	4473	4482
GAT	GCT	GAC	GGC	AAG	GAG
AGT	ACA	GCA	ATC	GAA	AAA
GTC	CTC	AAG	GCT	GAA	CCT
D	A	D	G	K	E
S	T	A	I	E	K
V	L	K	A	E	P
4491	4500	4509	4518	4527	4536
GAG	ATC	CTG	GAA	CTT	GAG
AGT	AAG	AGC	AGC	AAG	ATT
GTG	CTG	AAC	GTC	CTG	AAC
ATT	CAG	GTC	ATT	CAG	CAG
E	I	L	E	S	K
S	K	S	N	K	I
V	L	N	V	I	Q
4545	4554	4563	4572	4581	4590
ACA	GCC	GTT	GAC	CAG	TTC
GCA	CAG	ACA	CAG	TTC	GCA
CGT	ACA	GAA	ACA	GAA	ACA
GCC	CCC	ACT	CAT	GCT	TAT
T	A	V	D	Q	F
A	R	T	E	T	A
T	A	P	E	T	H
A	Y				
4599	4608	4617	4626	4635	4644
GAT	TCA	CAG	ACC	CAG	GTT
TGC	CCT	GCA	TGC	AGG	CTT
AGG	AGC	TGC	GAC	AGC	GAG
GAG	CCC	ACT	CAT	GAG	CCC
CCC	AAC	GAC	AGA	AGA	
D	S	Q	T	Q	V
P	A	C	R	L	D
L	D	S	R	E	P
N	R	P	N	R	
4653	4662	4671	4680	4689	4698
TGC	TGG	ACA	AAA	ATG	AAA
AGC	ACA	AAA	ATG	AAA	GAT
GCC	GCC	AAG	ATG	AAA	CAC
AGG	AGC	TGG	AGG	AGG	CCA
GCA	GTG	CGG	CGG	CCA	CCC
TGG	CCG	CAG	CAG	GTG	AGA
AGA	CCC				
C	W	T	K	M	K
D	A	K	M	K	H
V	P	A	K	H	P
P	V				
Q	P				
P	R				
R					
4707	4716	4725	4734	4743	4752
GAG	GAC	TTG	CAA	GTC	CTG
GAC	ACA	CAA	GTC	CTG	ACC
CTG	CTG	GTT	GAG	GCA	TGG
ACC	ACC	CTG	GAG	TGG	GCT
GCA	TGG	GCA	TGG	CAG	CCT
TGG	CGG	CAA	AAA	CAG	CGG
AGA	AAA	AAA	TGC	AAA	AAA
TGC	TGC	TGC	TGC	TGC	TGC
AGA	AGA	AGA	AGA	AGA	AGA
C	D	L	Q	V	L
E	P	R	K	R	C
R					
K					
C					

(30 of 90)

4761 4770 4779 4788 4797 4806
TTG CCG CGC TTG CAG TTG AAA GCG CCG GTG TCA AAG TAA GCA TTG AGA AGC TGC

L P R L Q L K A P V S K *

4815 4824 4833 4842 4851 4860
CTC CTC AAC CCA AAG ATC CAA AAG GAG CAT GCT GCT GAT GGC CCT CAG CTC CAA

4869 4878 4887 4896 4905 4914
AGC TTA GCC CAG GCA GAG GCC AGT GCC TCT GGA AAC CTA ACC AAA GAA TCC CCA

4923 4932 4941 4950 4959 4968
GAC ACC ACC GGA CCA AAG CTA ACC GAG GAG GGC GAT CCC CCA AAA GTT CAG GTC

4977 4986 4995 5004 5013 5022
CAG GAA GAA GAA ATG TCT ACC AAG TCA GTC AAA GAG AAC AAG GCC CAG GCA GAA

5031 5040 5049 5058 5067 5076
GAG GAC CTG CAG GAG CCA AAG GGA GAC CTG GCA GAA TCC TCC GAT GTT AGT TGC

5085 5094 5103 5112 5121 5130
TCA TTG TAC ATC TGT AAG ACC AGA ATG TGA AAA CAA GTC ACA GAA CAA GAT GCT

5139 5148 5157 5166 5175 5184
GCT GTT GGG ACC TTG AGA CCA AGA TTT CAG AGC CCA TGA CAT CCA GAG AGC AGG

5193
GCC GTC CAA TGA TTT C 3'

FIG.11L

(31 of 90)

SSeCKS
13.2.2

220 KD -



116 -

97.4 -

FIG.12

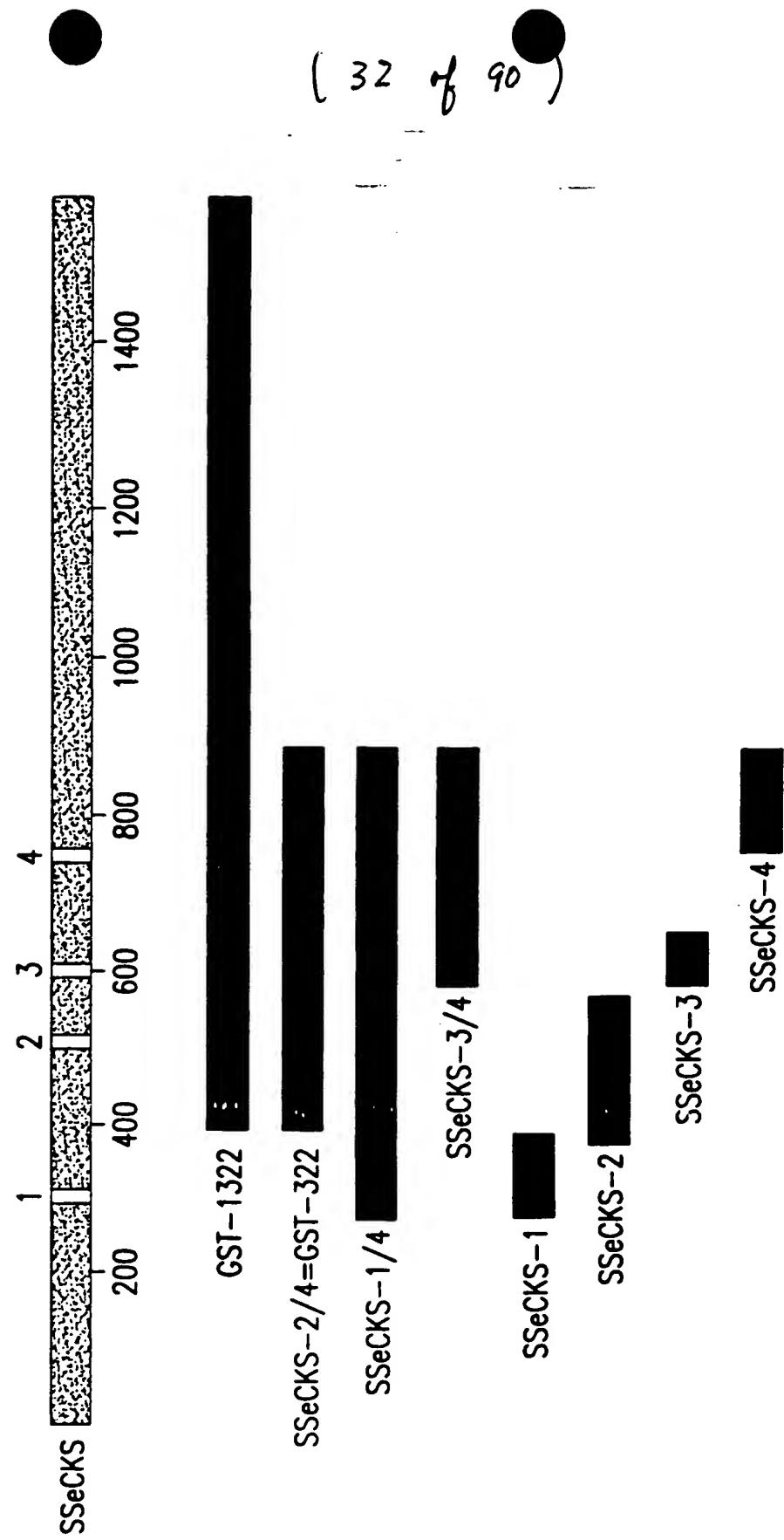


FIG. 13A

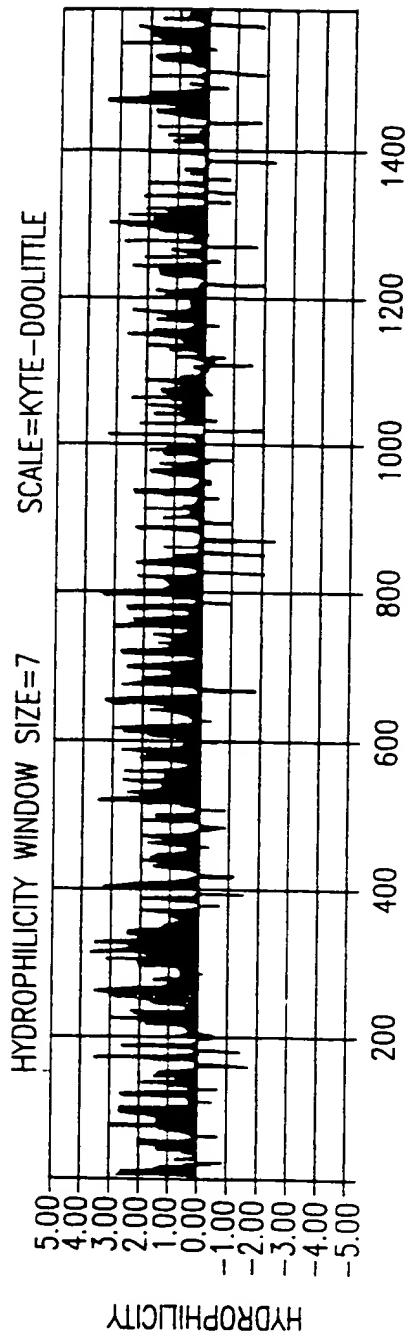


FIG. 13B

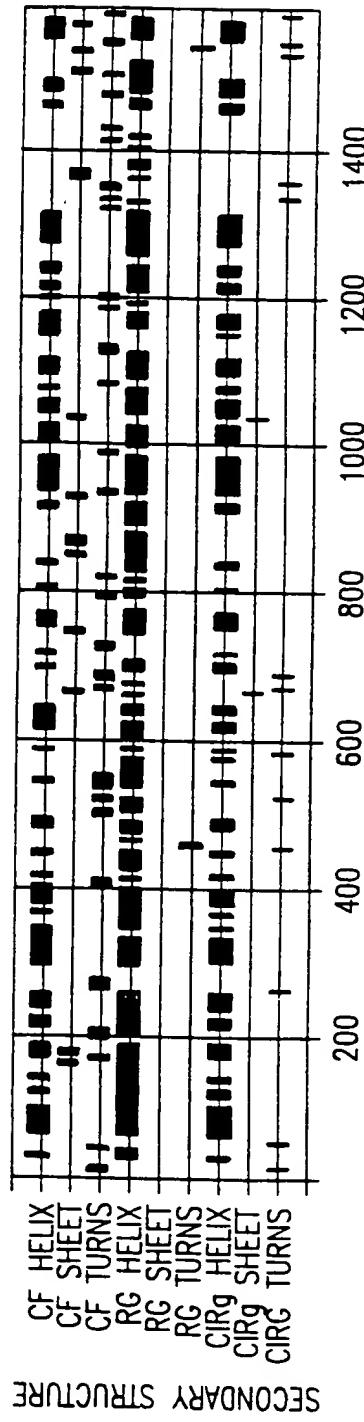


FIG. 13C

(34 of 90)

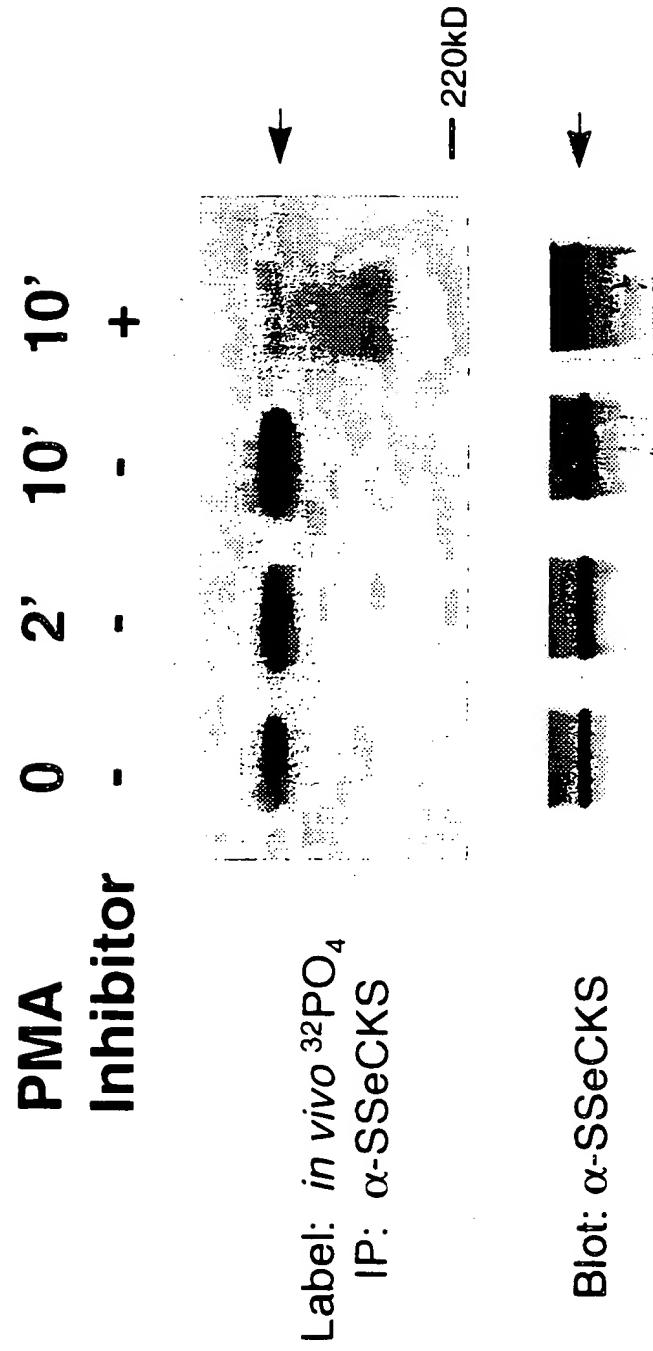


FIG. 14

(-35 of 90)



FIG. 15A

FIG. 15B

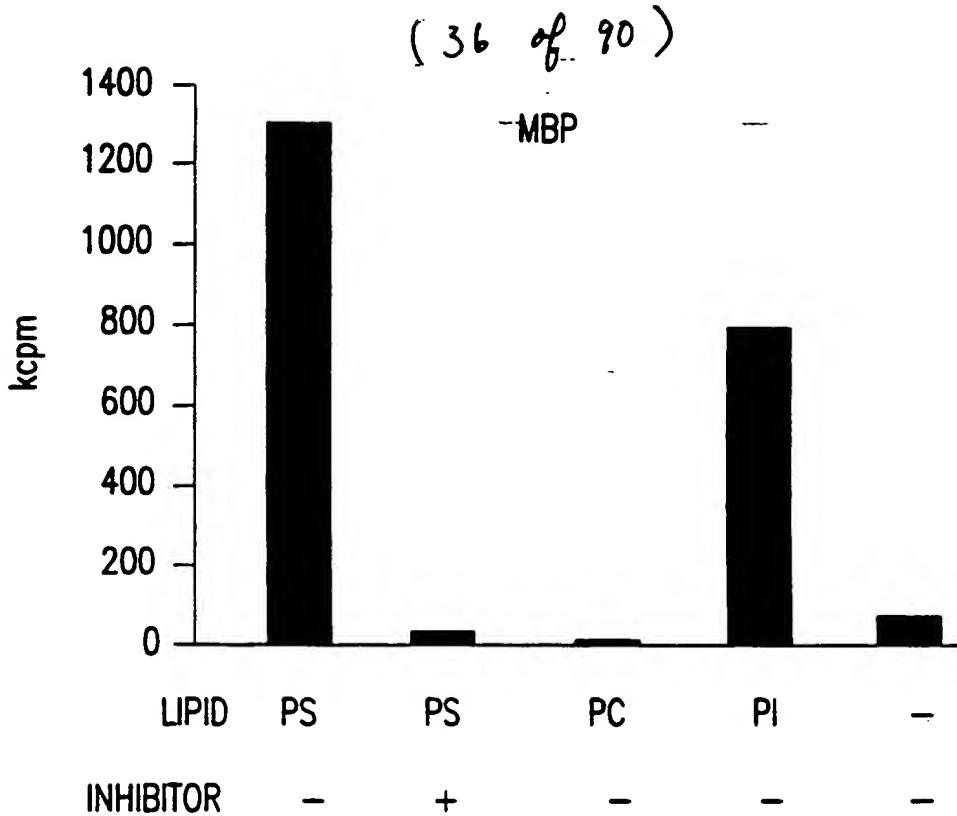


FIG.16A

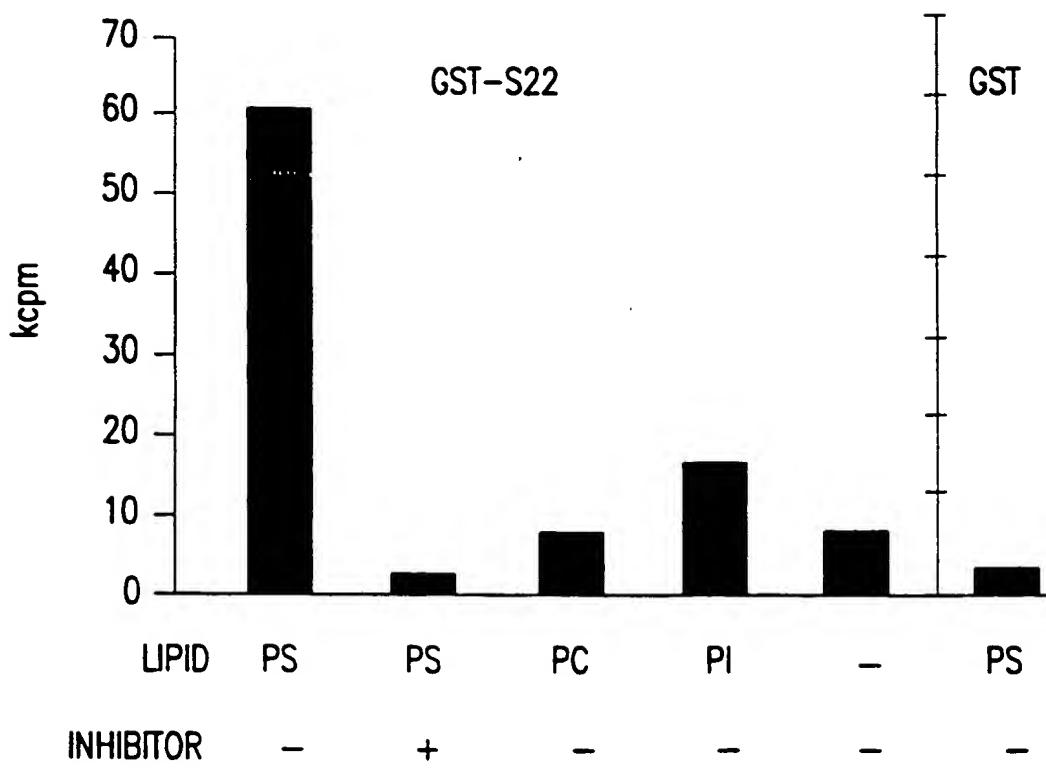


FIG.16B

(37 of 90)

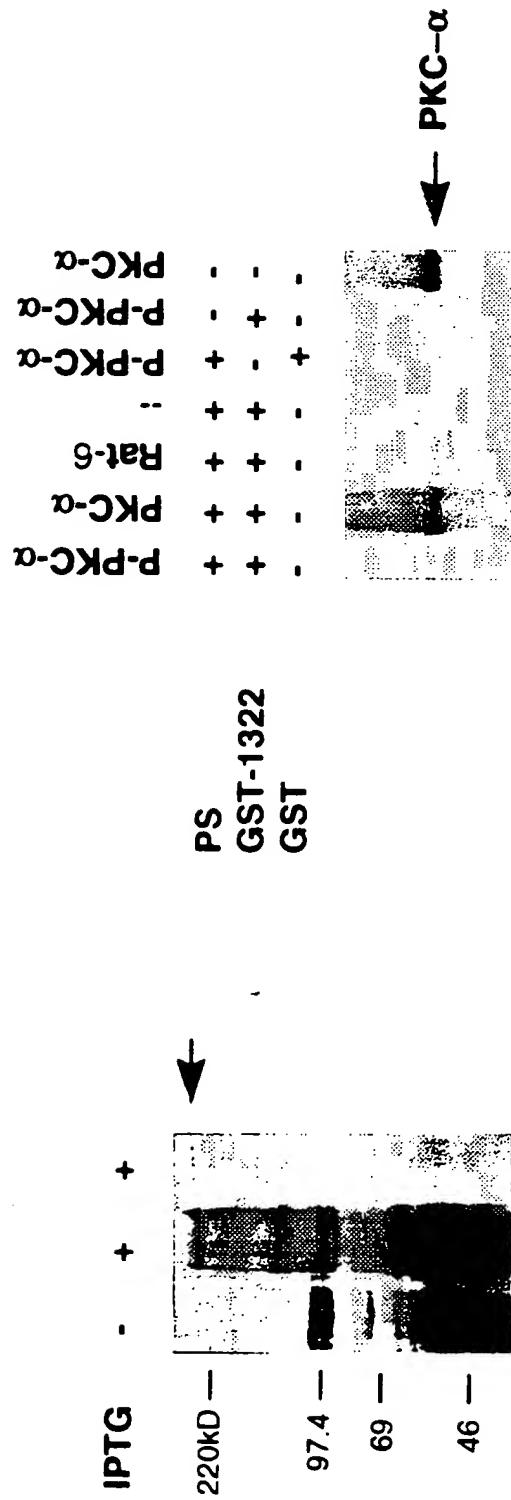


FIG. 17A

FIG. 17B

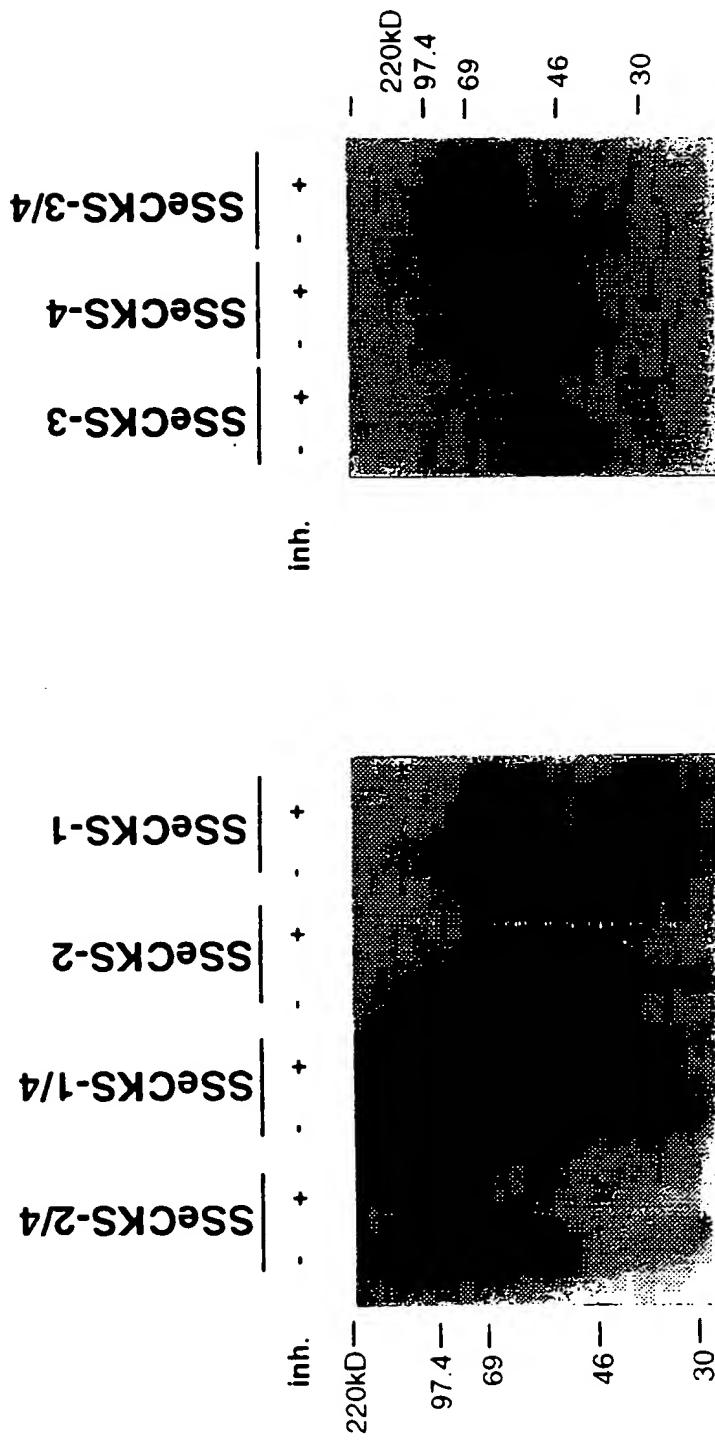


FIG. 18A

FIG. 18B

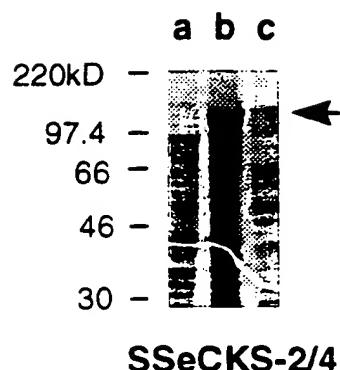


FIG.18C

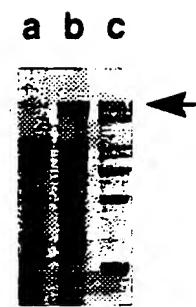


FIG.18D

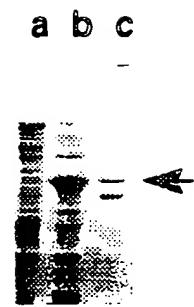


FIG.18E

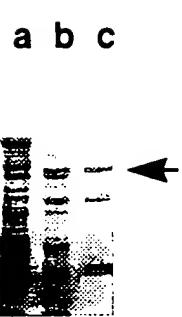


FIG.18F

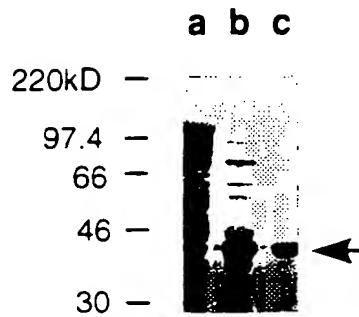


FIG.18G

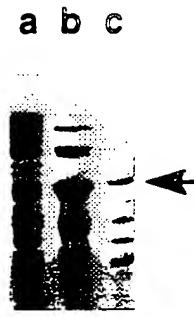


FIG.18H



SSeCKS-3/4

FIG.18I

(40 of 90)

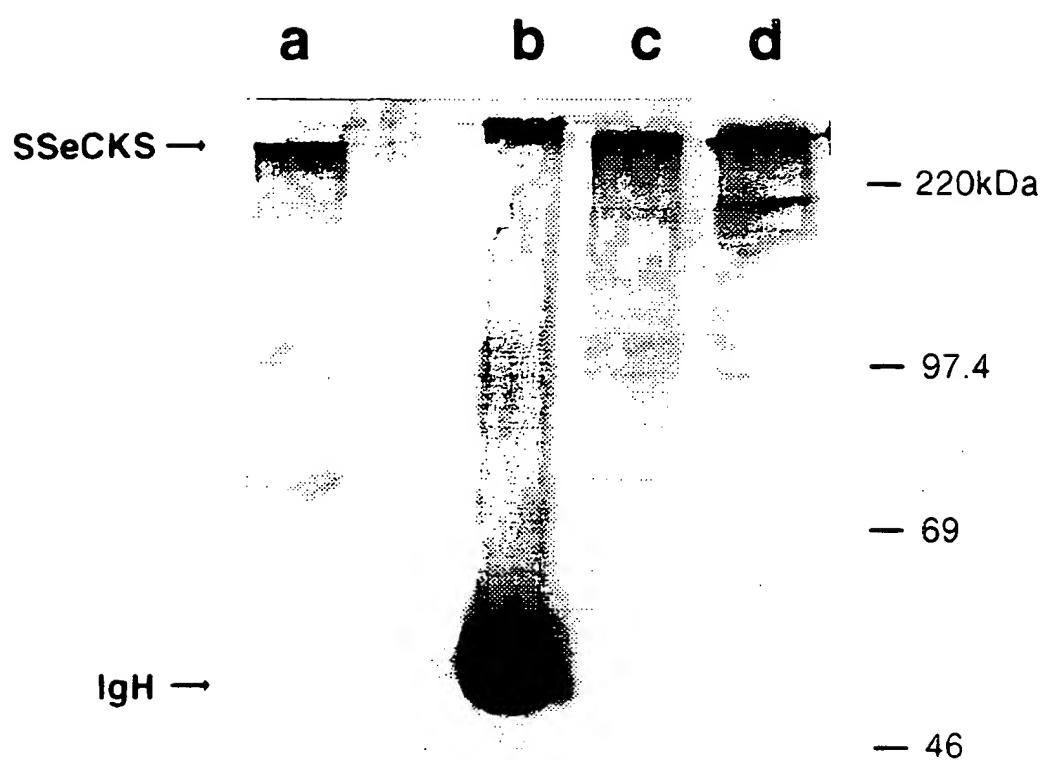


FIG. 19

(41 of 90)

220KD —

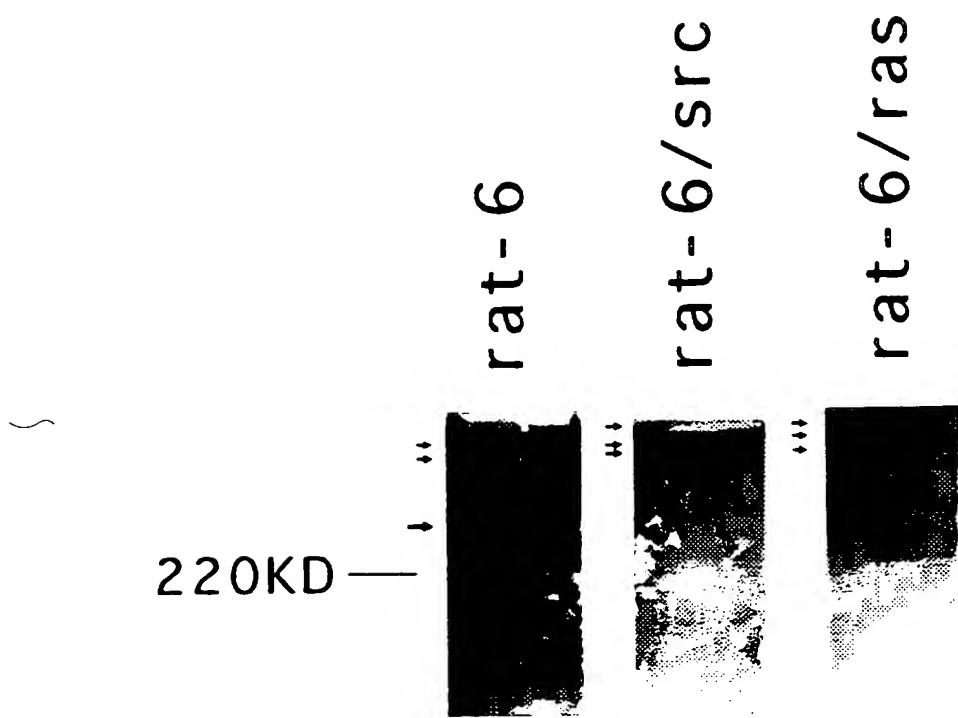


FIG.20

(42 of 90)



FIG.21A



FIG.21B



FIG.21C



FIG.21D



FIG.21E

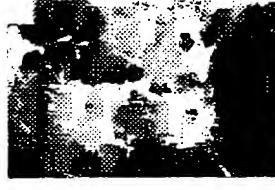


FIG.21F

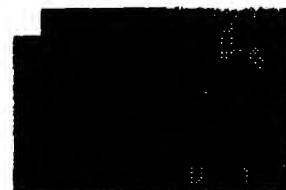


FIG.21G



FIG.21H



FIG.21I



FIG.21J

(43 of 90)

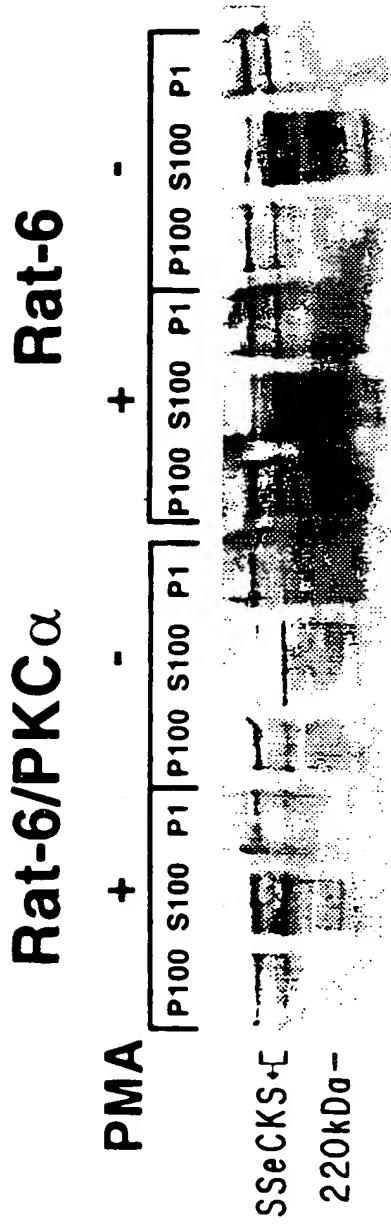


FIG. 22

(44 ~~of~~ 90)

spleen
thymus
prostate
testes
ovary
small intestine
colon
PBL



FIG. 23A

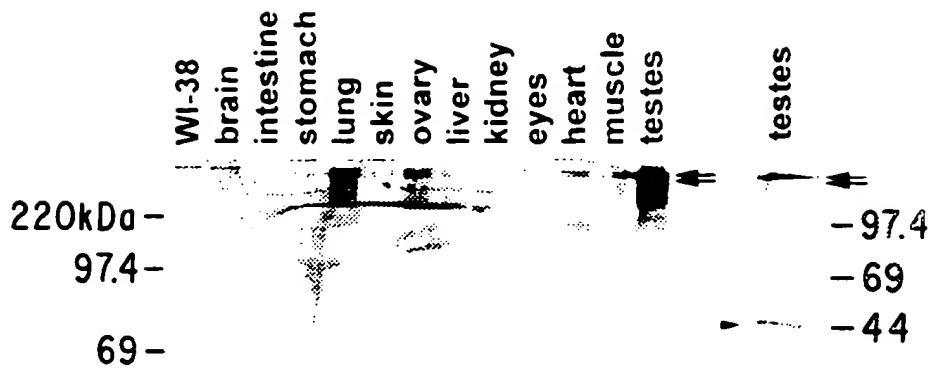


FIG. 23B

(45 ° 90 °)

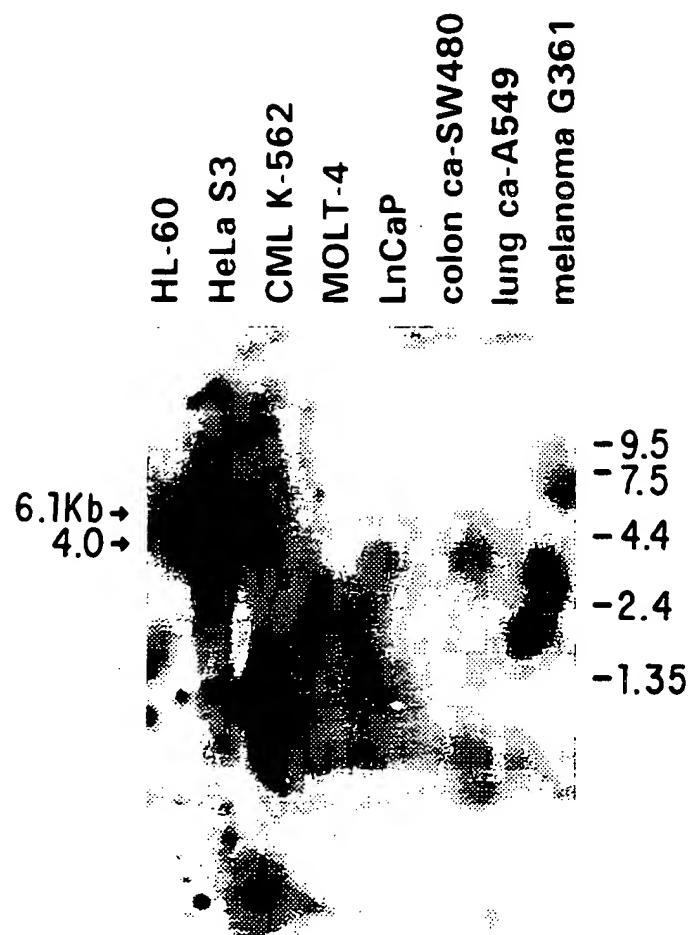


FIG. 24

(46 of 90)

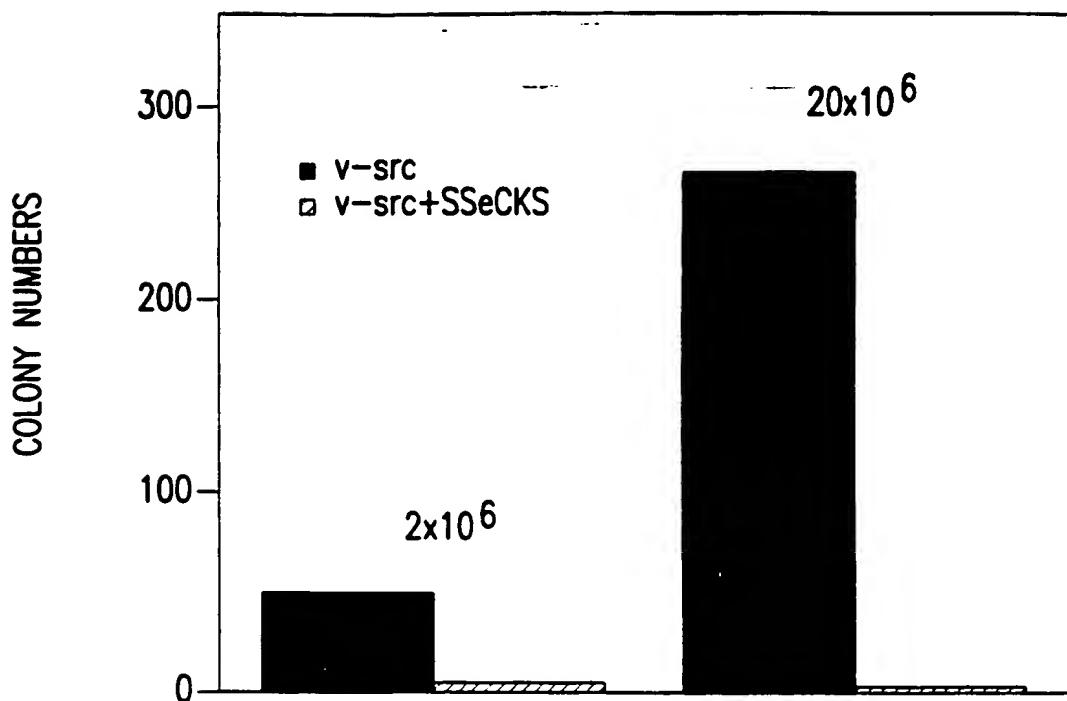


FIG.25A

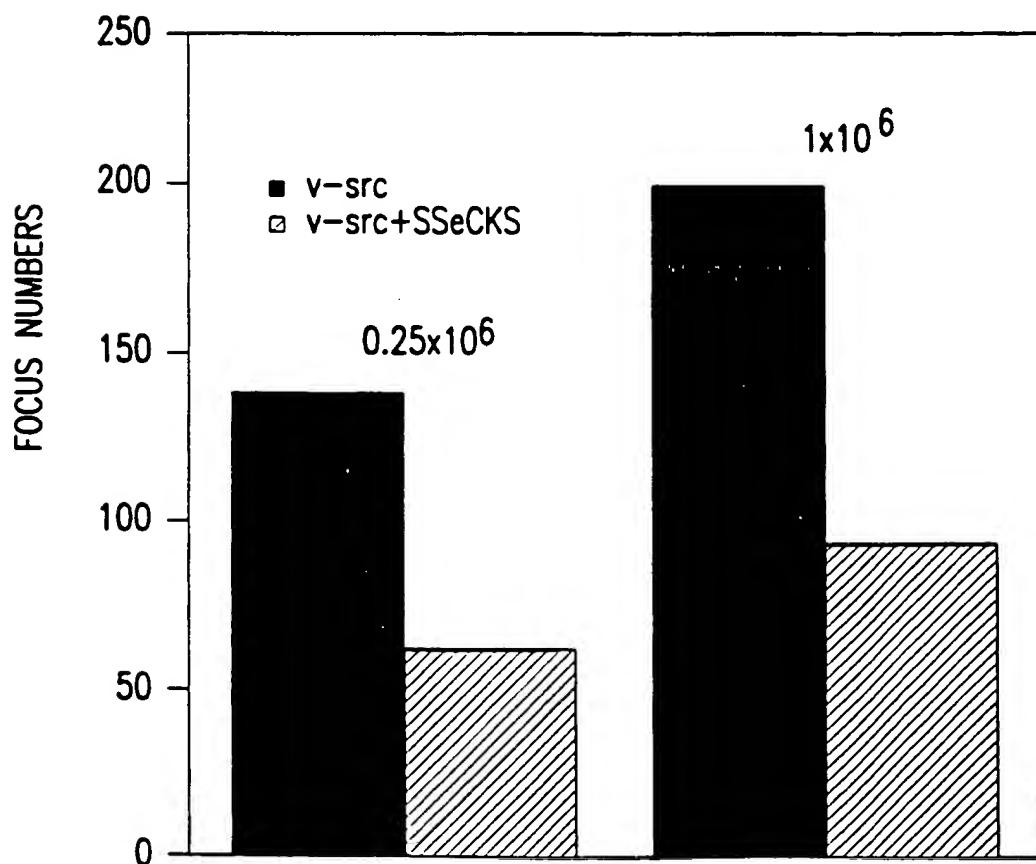


FIG.25B

(47 of 90)

		<u>Myr.</u>	<u>Pal.</u>
src	MGSSKSKPKD	+	-
yes	MGCIKSKEDK	+	+
SSeCKS	MGAGSSTEQR	+	?
G _α t1	MGAGASAEEK	+	-
G _α i1	MGCTLSAEDK	+	+
GAP-43	MLCCMRRTKQ	-	+
MYRIST. CONCENSUS:	MGXXX _T ^S		

FIG.26

(48 of 90)

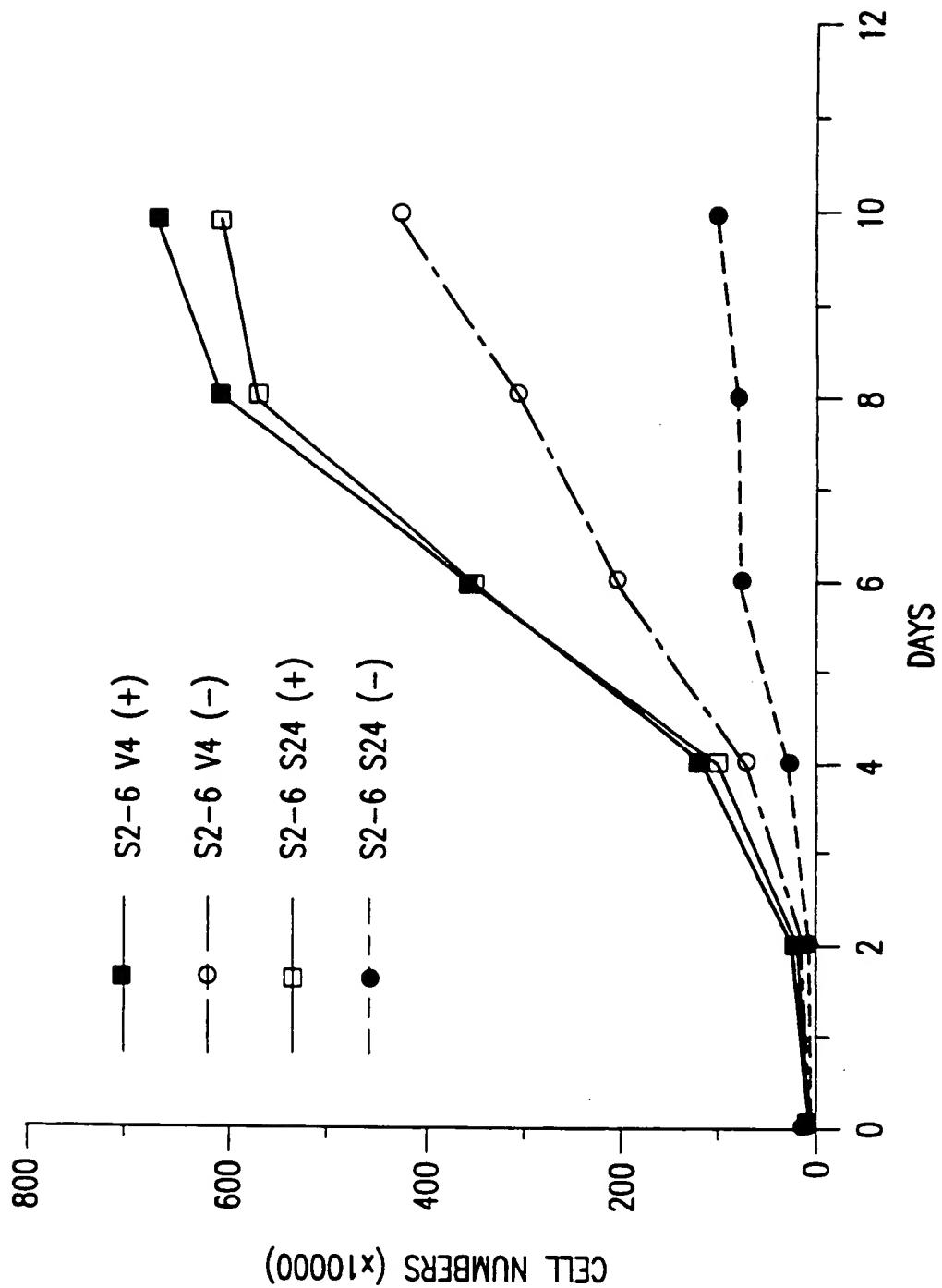


FIG.27

(49 of 90)

Tet + -



- 220kDa

FIG.28

(50 of 90)

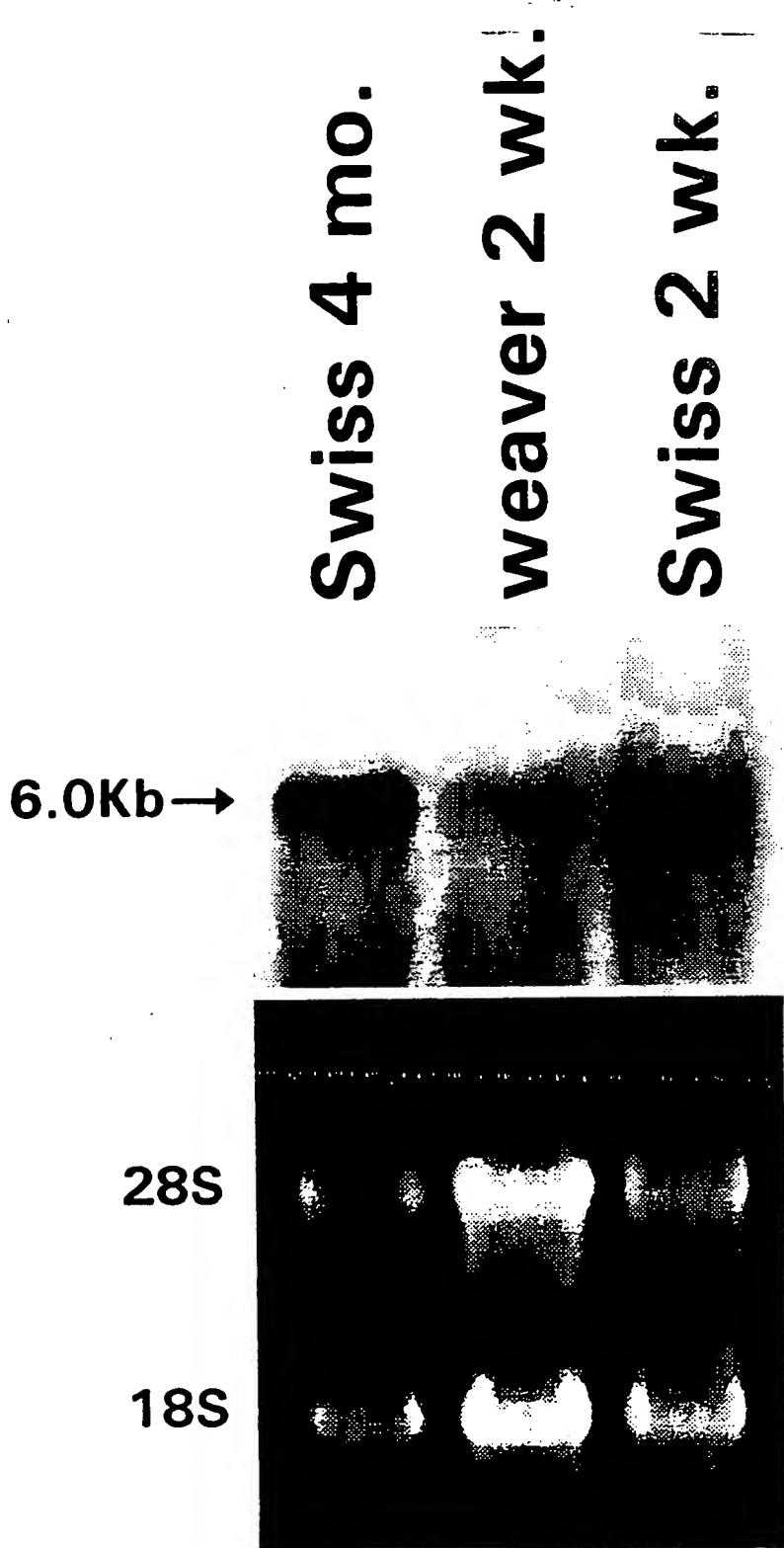


FIG. 29

(51 of 90)

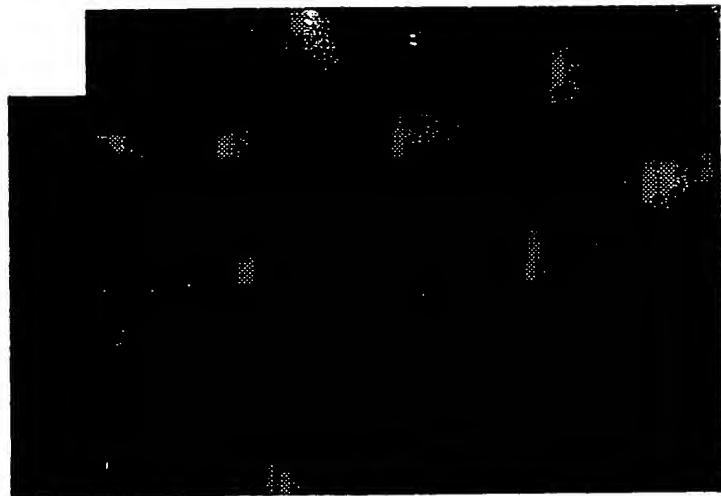


FIG.30A



FIG.30B

(52 of 90)

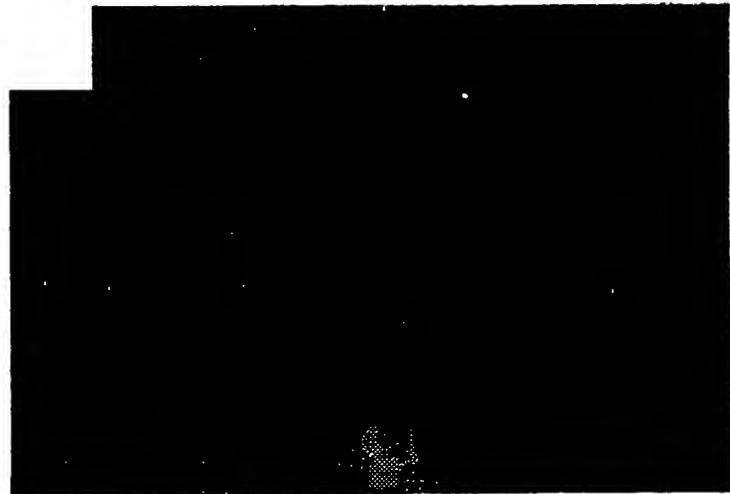


FIG.30C

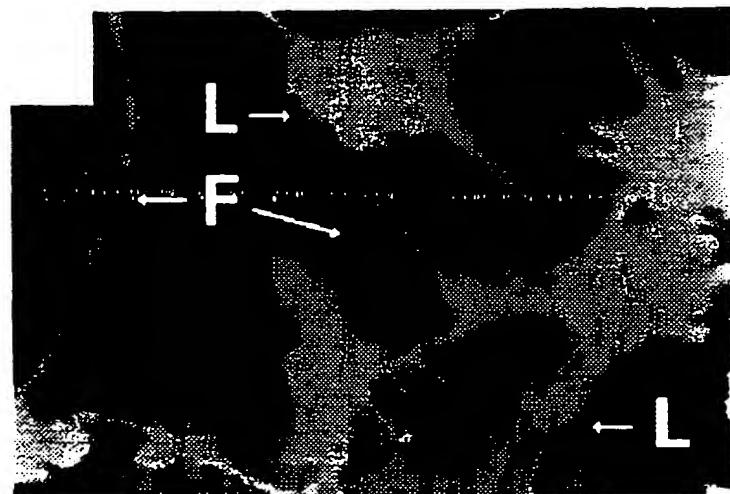


FIG.30D

(53 of 90)

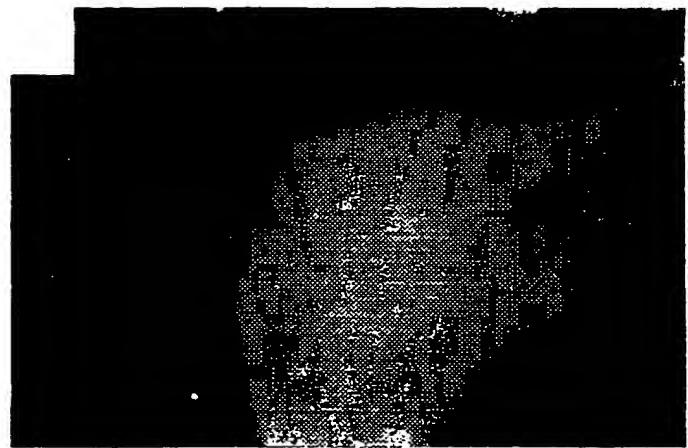


FIG.31A



FIG.31B

(54 of 90)



FIG.31C



FIG.31D

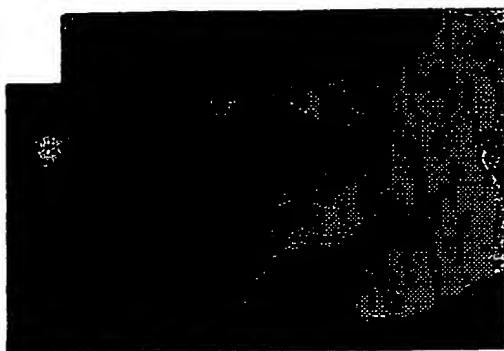


FIG.32A

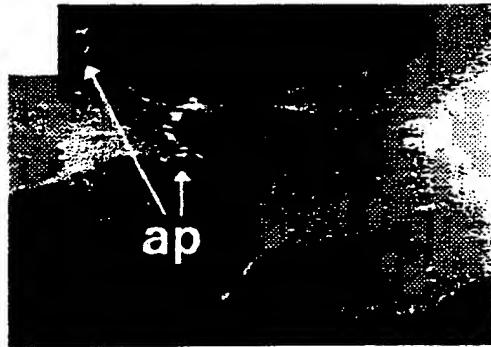


FIG.32B



FIG.32C

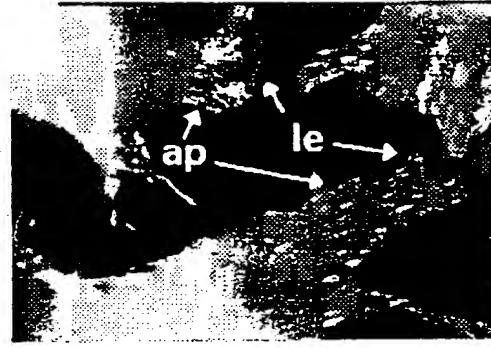


FIG.32D



FIG.32E



FIG.32F

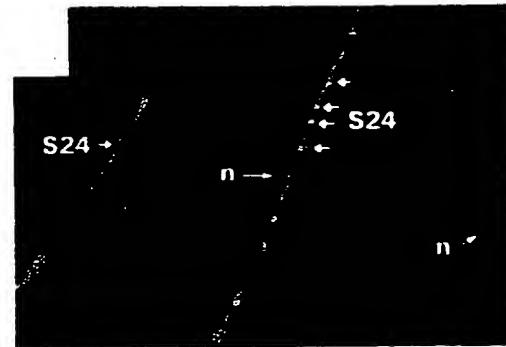


FIG.32G

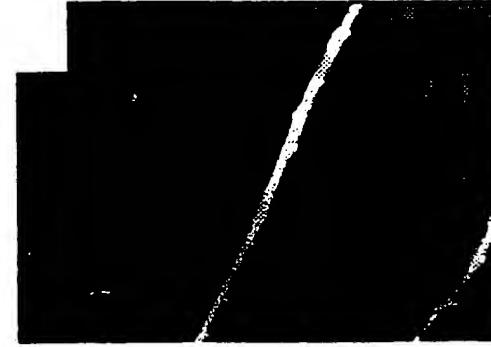


FIG.32H

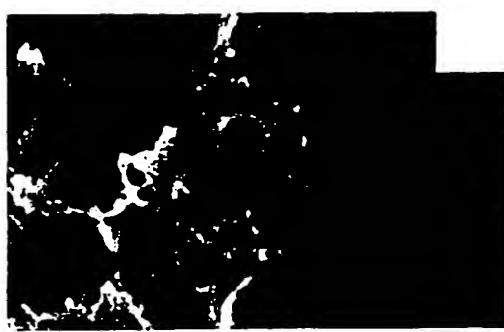


FIG.33A



FIG.33B



FIG.33C



FIG.33D



FIG.33E



FIG.33F



FIG.33G



FIG.33H

(57 of 90)

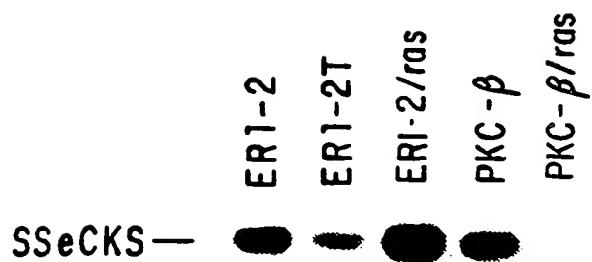


FIG.34

(58 of 90)

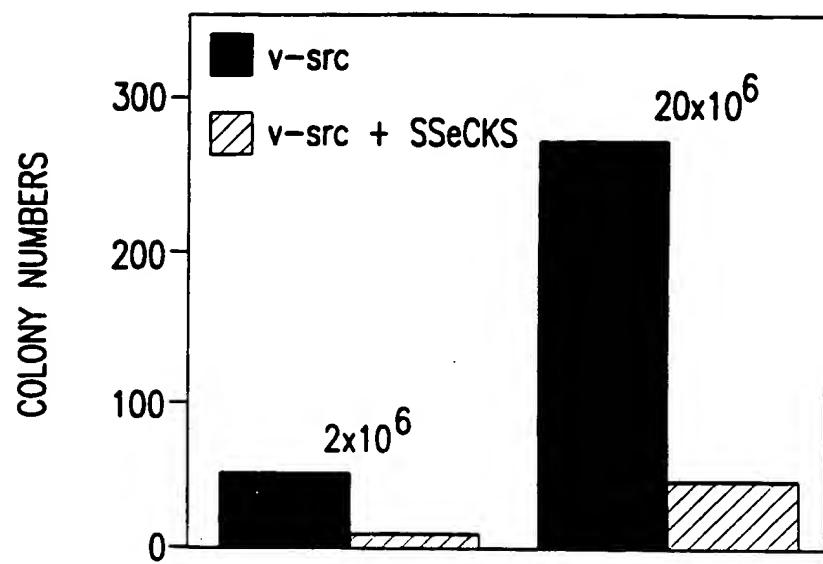


FIG.35A

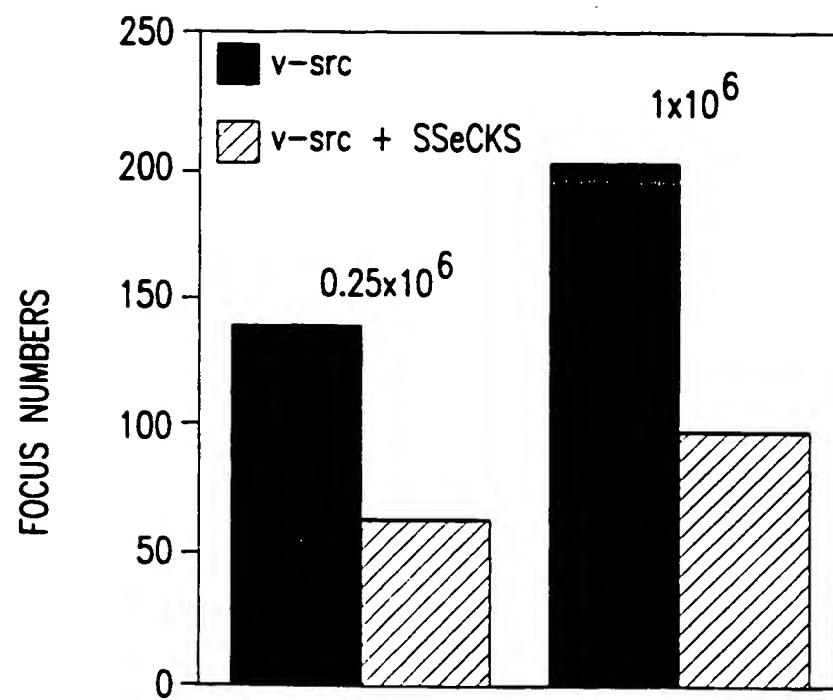


FIG.35B

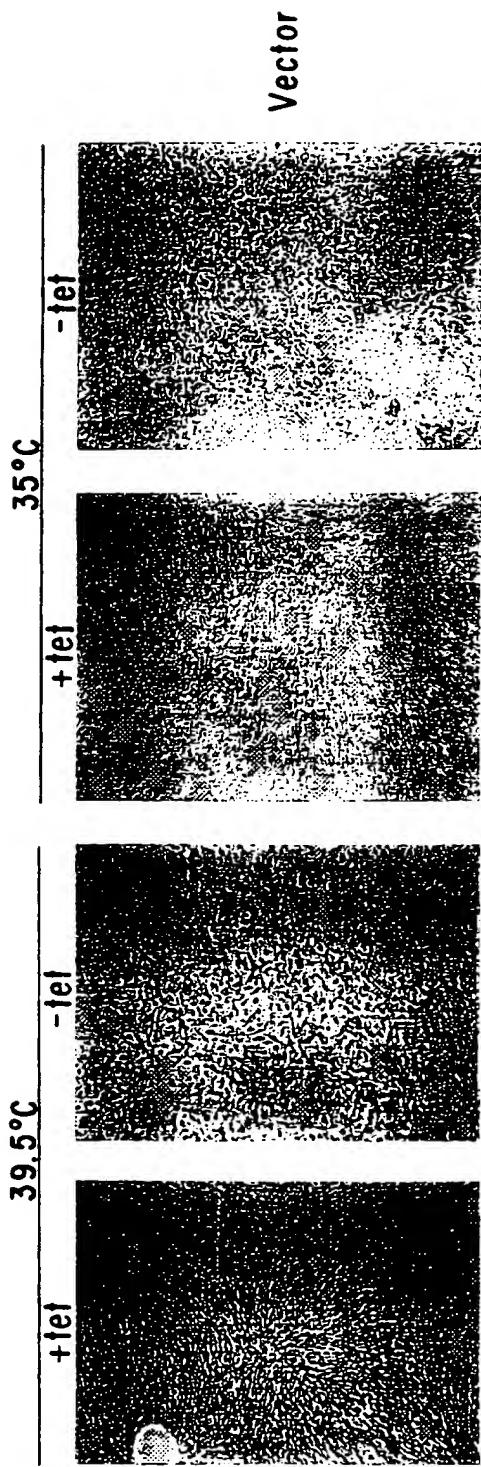


FIG. 36A-1 FIG. 36A-2 FIG. 36A-3 FIG. 36A-4

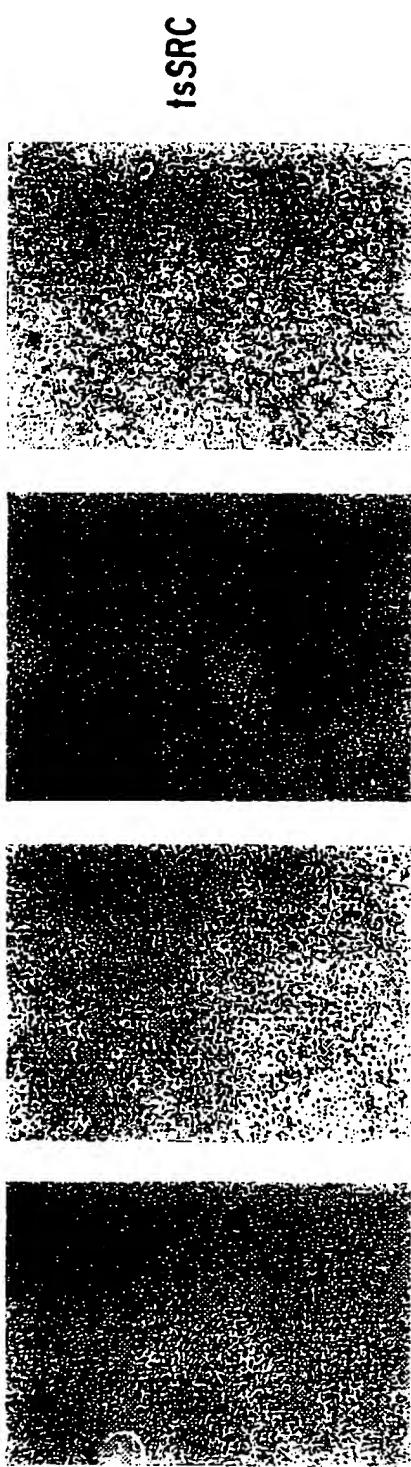


FIG. 36A-5 FIG. 36A-6 FIG. 36A-7 FIG. 36A-8

(60 of 90)

35°C



FIG.36B-1

39.5°C

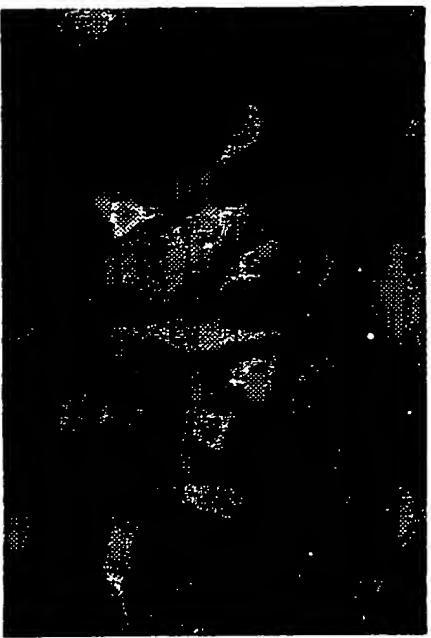


FIG.36B-2

+161

-161

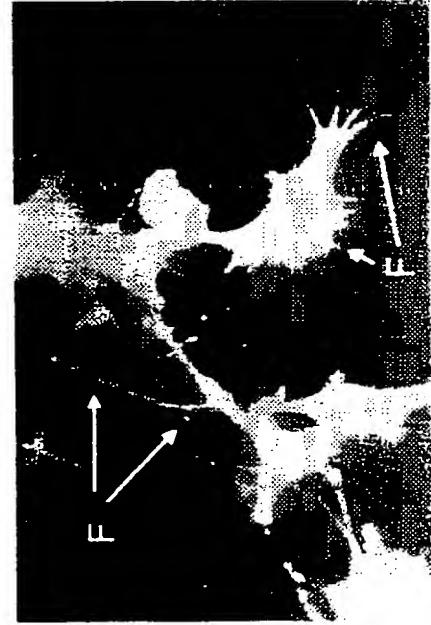


FIG.36B-3

FIG.36B-4

(61 of 90)

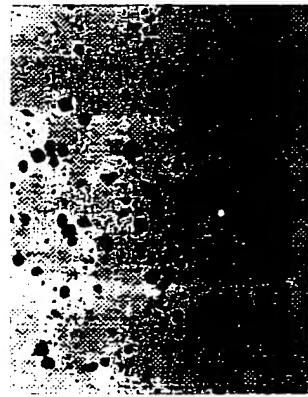


FIG.37A-1



FIG.37A-2

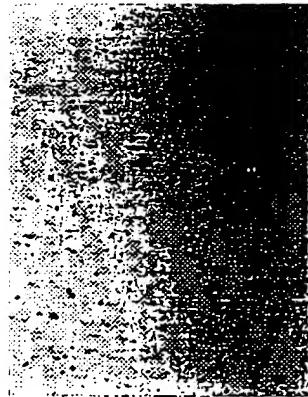


FIG.37A-3

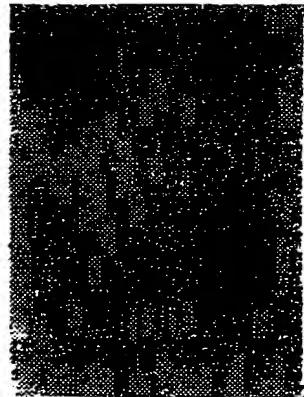


FIG.37A-4

(62 of 90)

SOFT AGAR COLONY FORMATION						
	ts src1	ts src2	ts src3	ts src4	pLJ2	pLJ3
+ tet	2160	1640	2800	1080	0	0
- tet	60	60	110	35	0	0

FIG.37B

(63 of 90)

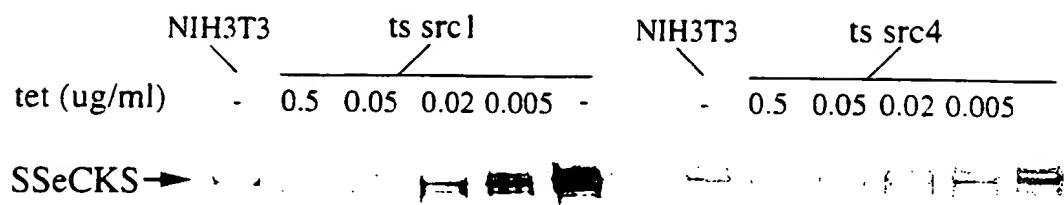


FIG.38A

0.5ug/ml tet



FIG.38C-1

0.02ug/ml tet

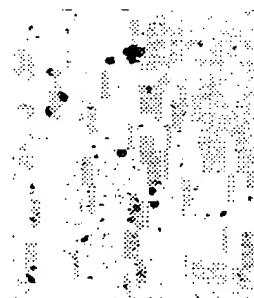


FIG.38C-2

(64 of 90)

tet (ug/ml)	SOFT AGAR COLONY FORMATION				39°C
	35°C				
ts src1	2852	2464	174	51	22
ts src4	1463	743	67	11	0

FIG. 38B

(65 of 90)

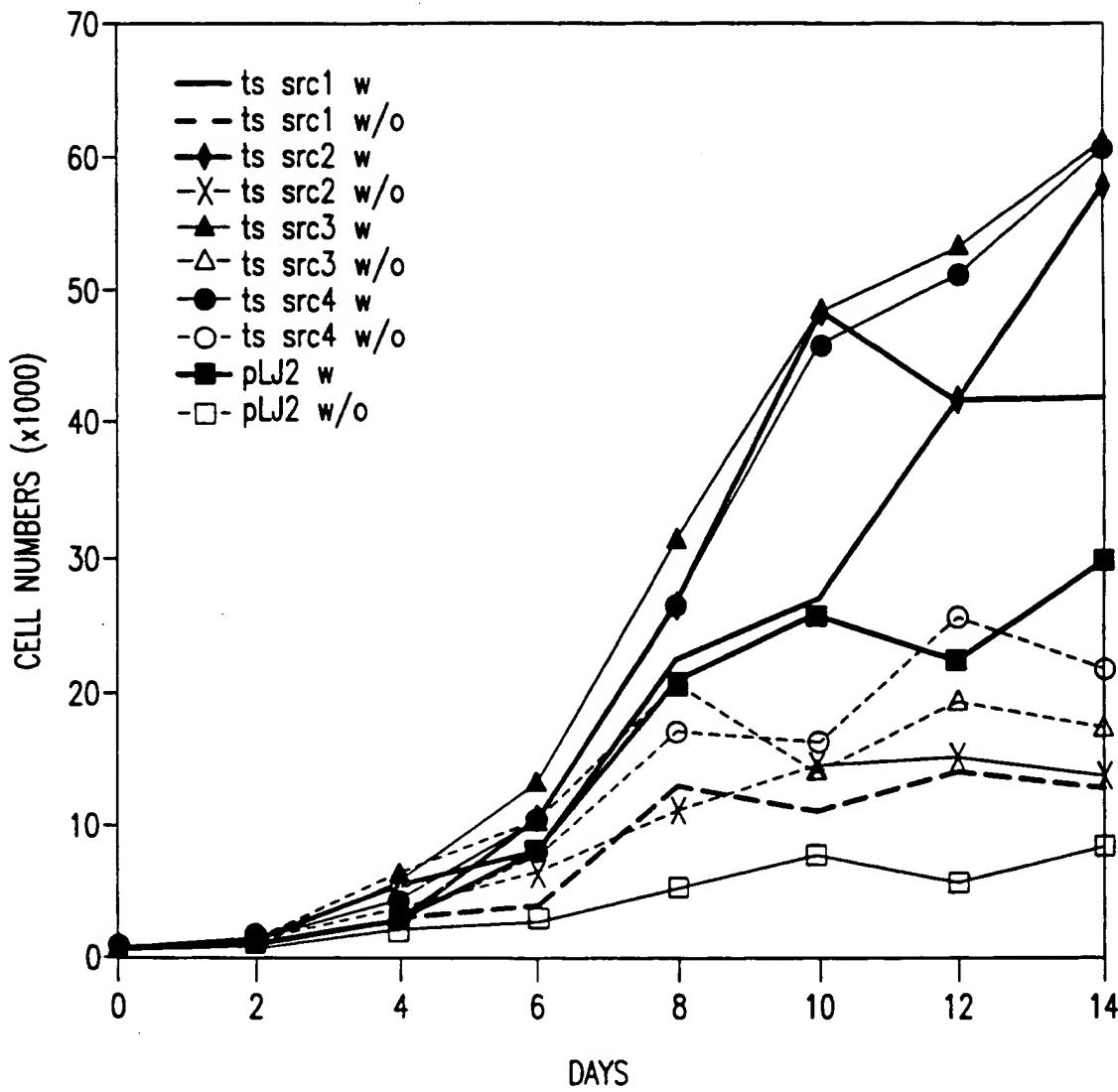


FIG.39A

(66 of 90)

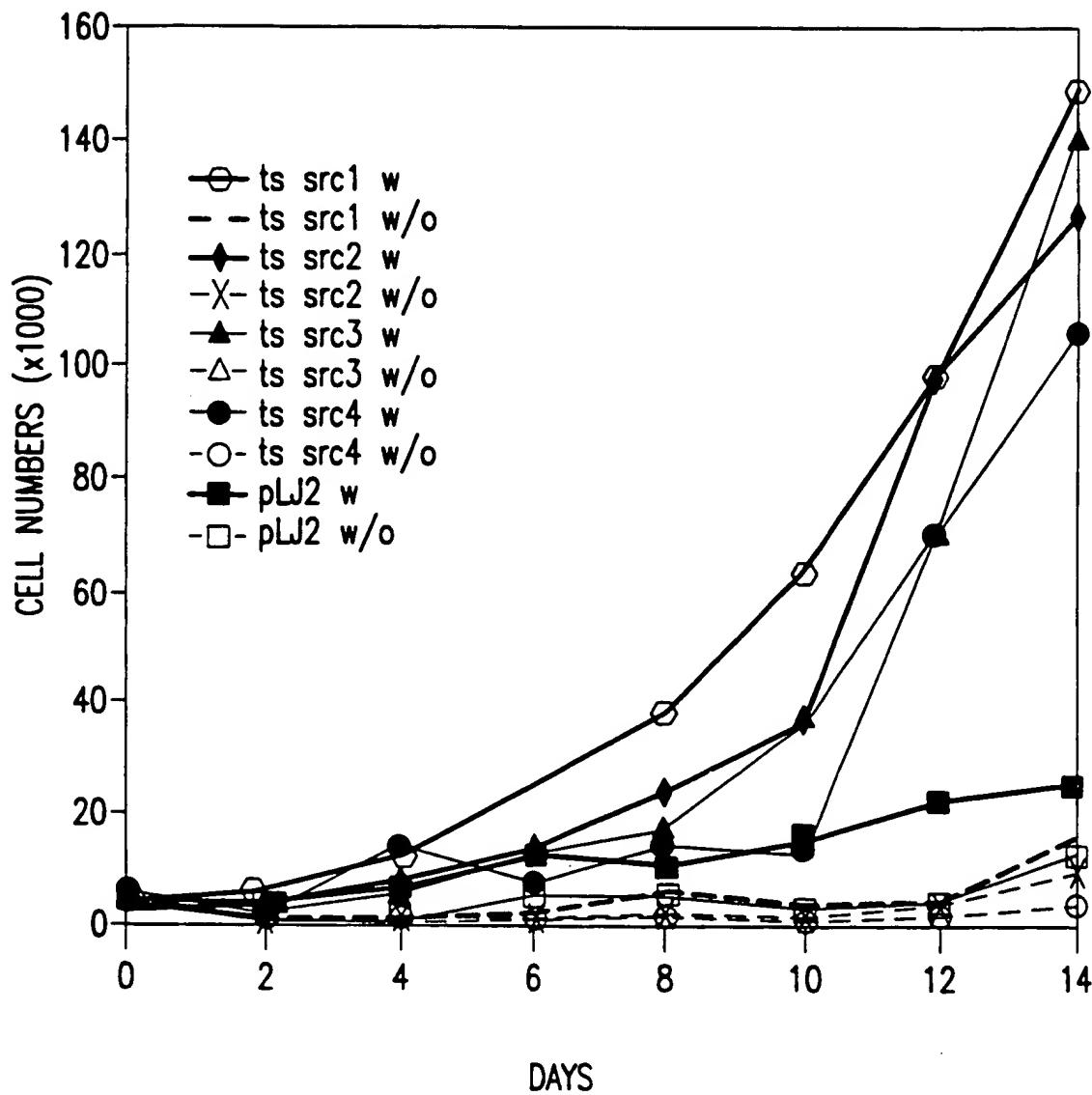


FIG.39B

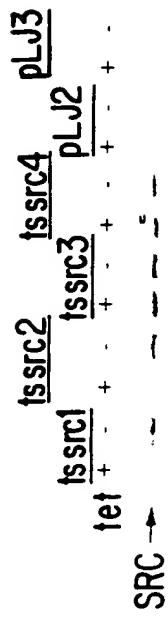
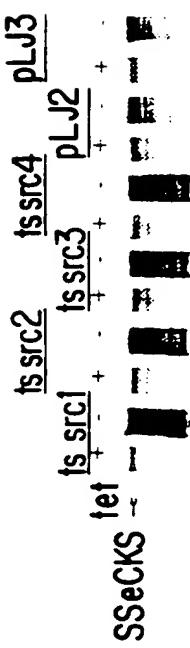


FIG. 40B

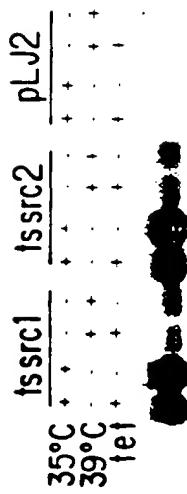


FIG. 40C-1

FIG. 40C-2

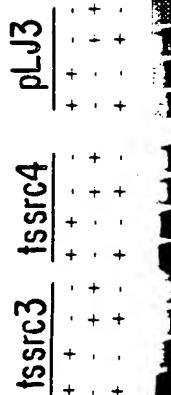


FIG. 40D-1

FIG. 40D-2

(67 of 90)

(68 of 90)

	fs src1	fs src2	plj2	fs src3	fs src4	plj3
35°C	+	-	+	-	+	-
39°C	-	+	-	+	-	+
tet	+	-	+	-	+	-

FIG. 4.1 A-1

FIG. 41 A-2

	35°C	39°C	41°C
ts src 1	+	-	-
ts src 2	+	+	-
plj 2	+	+	-
ts src 3	+	+	-
ts src 4	+	+	-
plj 3	+	+	-

FIG. 41B-1

FIG. 41B-2

ts src4
ts src3
ts src1 ts src4 plj2 plj3

FIG. 41 C

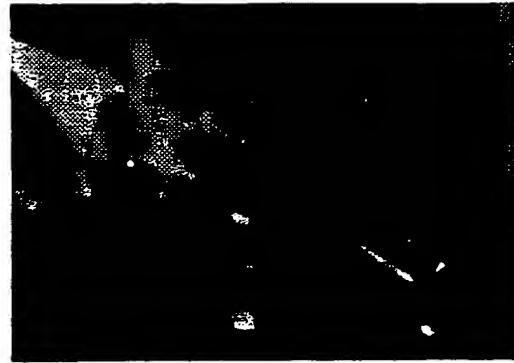
(69 of 90)

SSeCKS



FIG.42A-1

Vinculin



35°C
+tet

FIG.42A-2



FIG.42A-3



35°C
-tet

FIG.42A-4

(70 of 90)

SSeCKS



FIG.42A-5

Vinculin



39.5°C
+tet

FIG.42A-6



FIG.42A-7



39.5°C
-tet

FIG.42A-8

(71 of 90)

SSeCKS



FIG.42B-1

Phalloidin



35°C
+ tet

FIG.42B-2



FIG.42B-3



35°C
- tet

FIG.42B-4

(72 of 90)

SSeCKS



FIG.42B-5

Phalloidin



FIG.42B-6



FIG.42B-7



FIG.42B-8

42072000-00000000

Figure 43

(73 of 90)

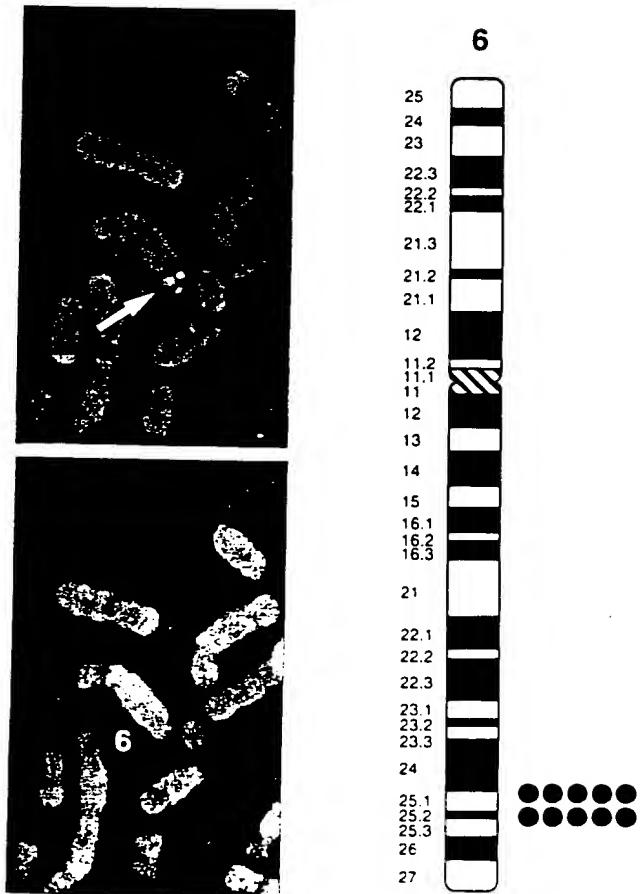


Figure 44

(74 of 90)

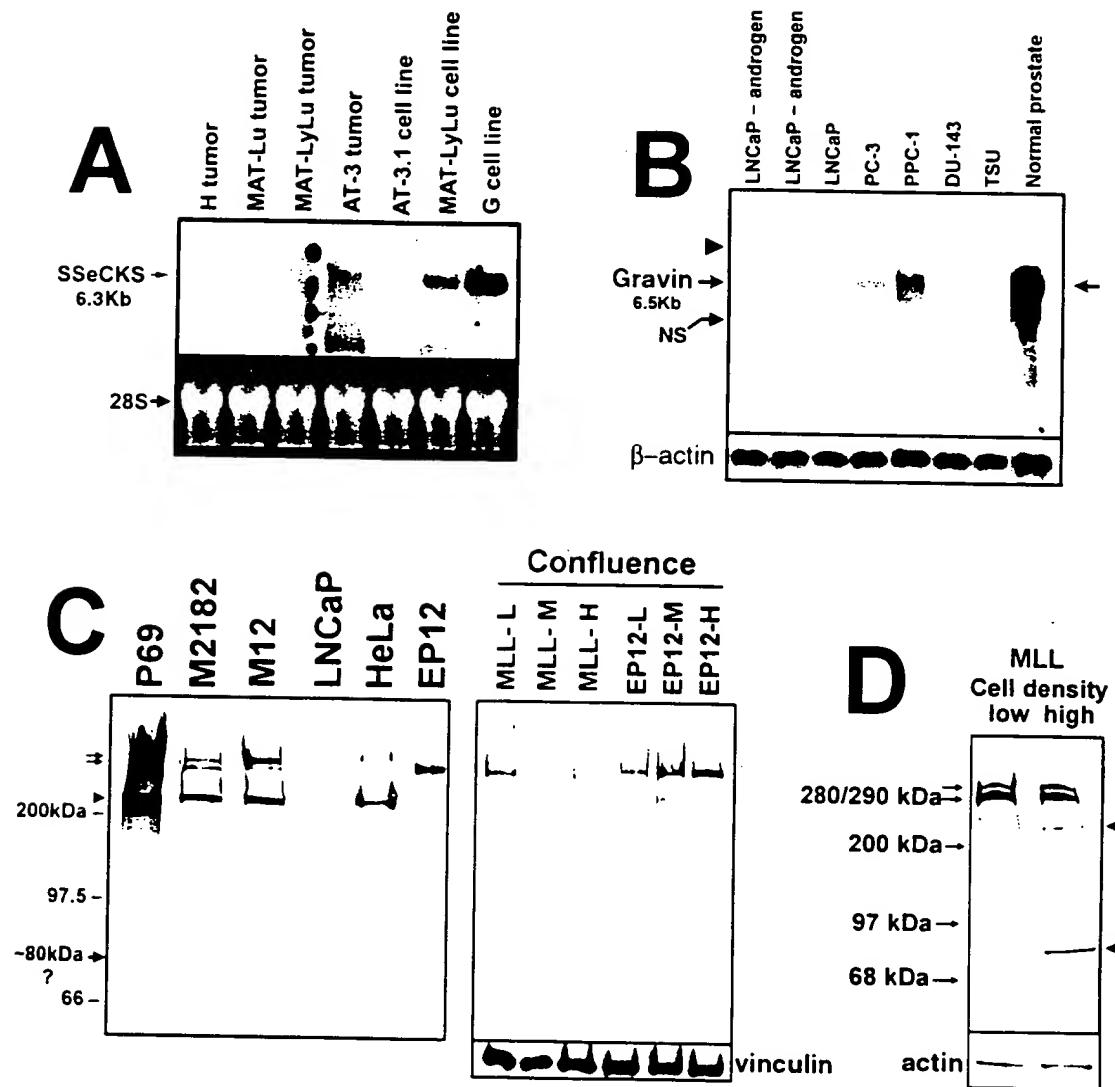


Figure 45

(75 of 90)

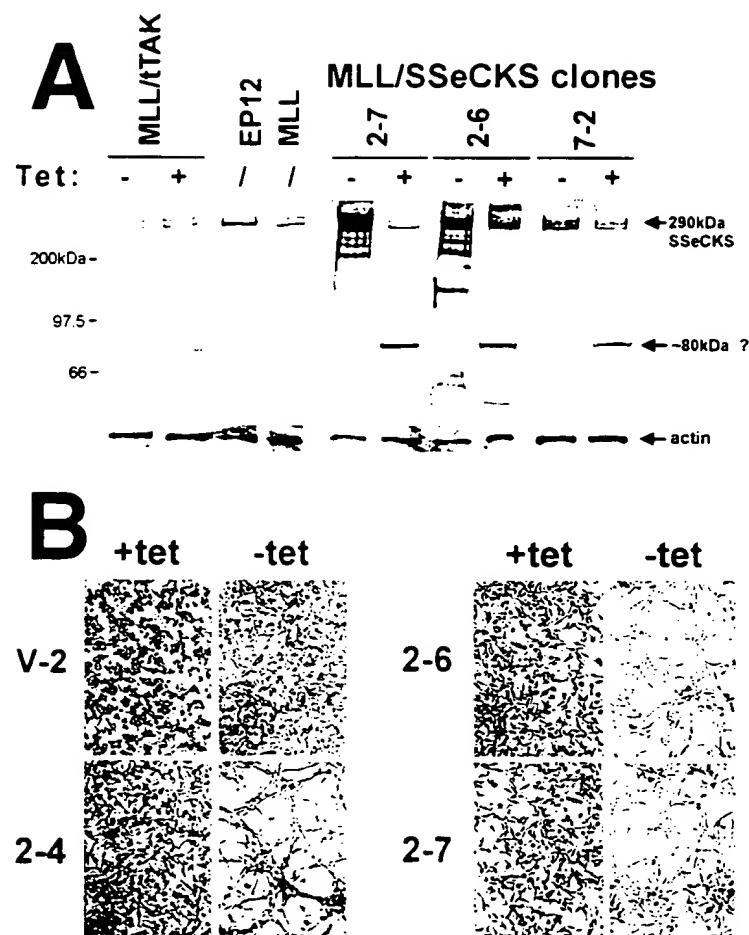
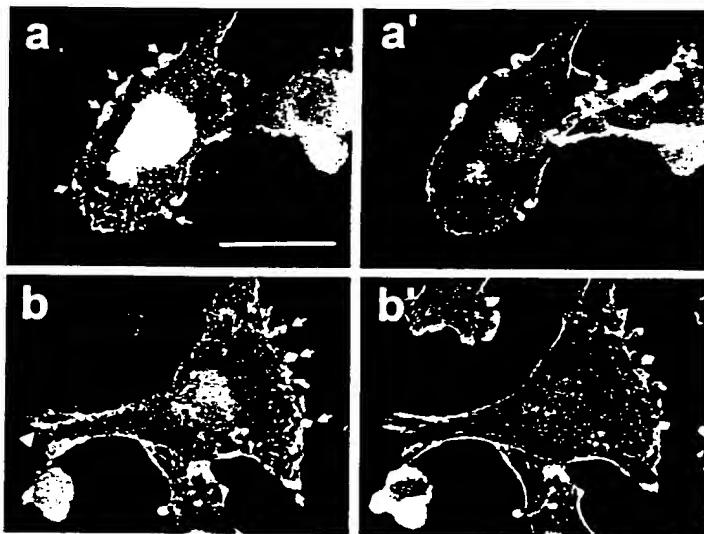


Figure 4b

(7b of 90)

A



B

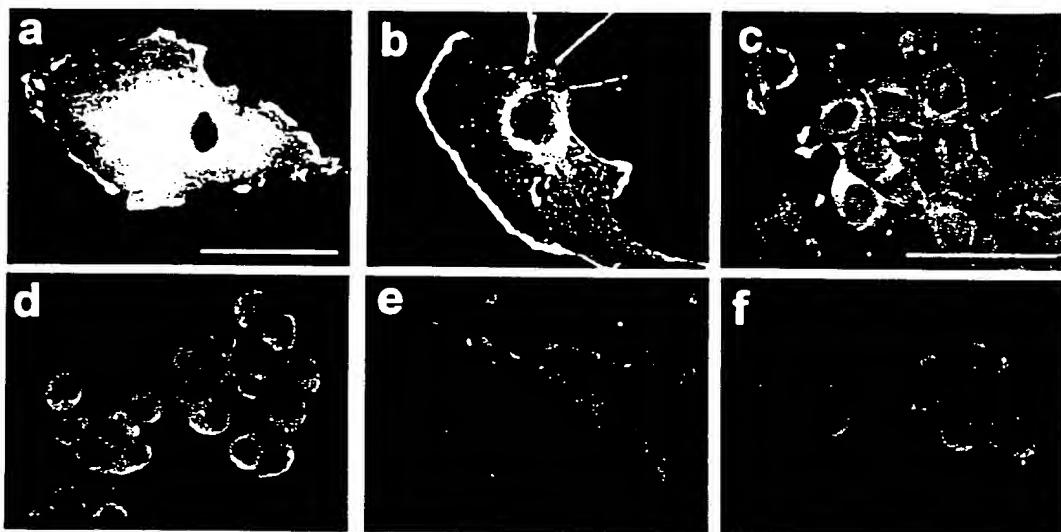
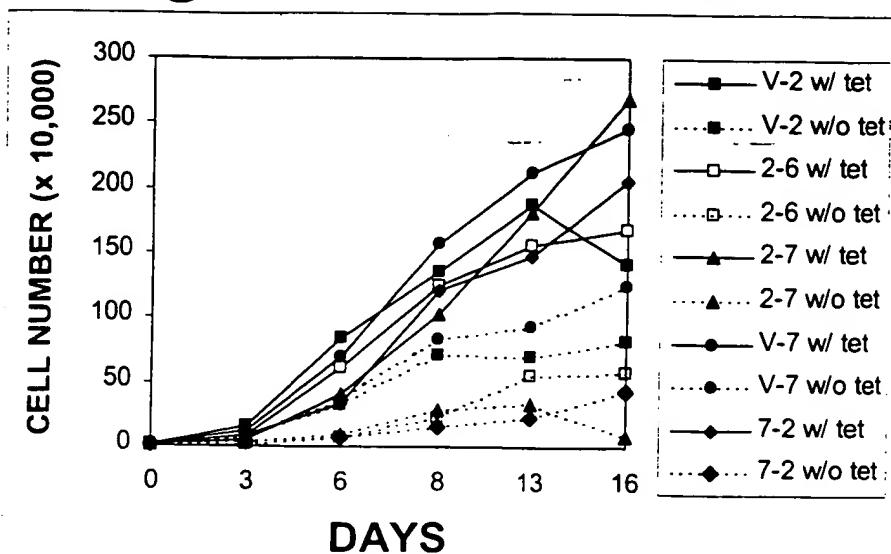
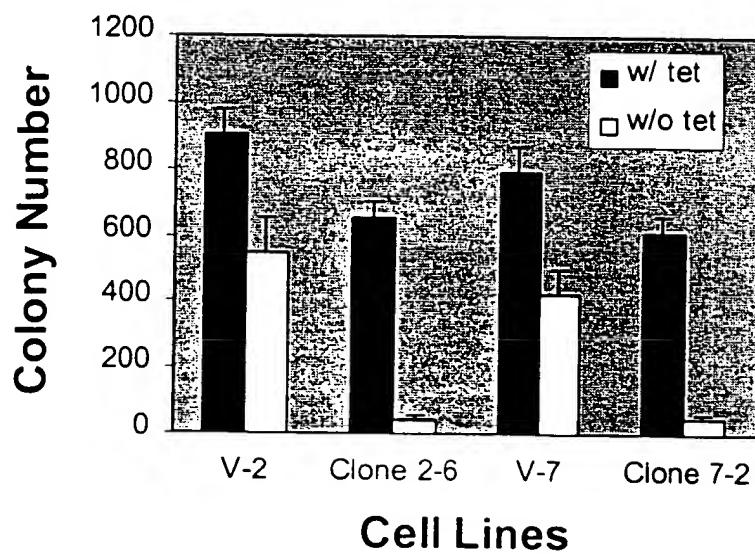


Figure 47 (77 of 90)

A



B



C

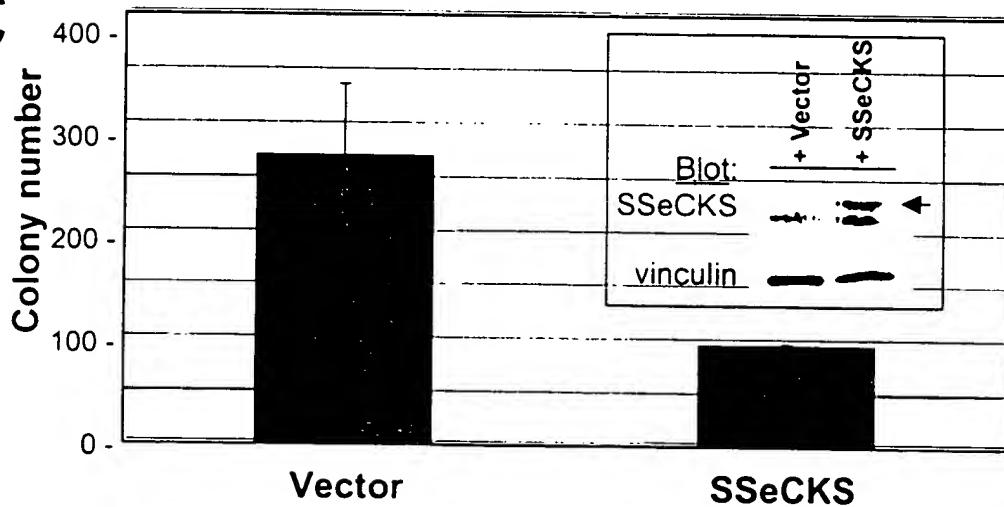
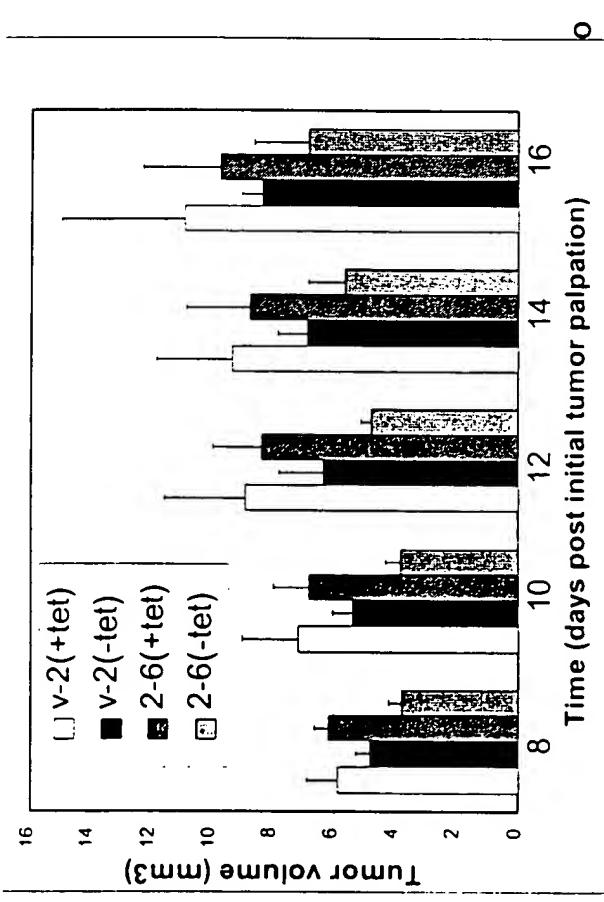
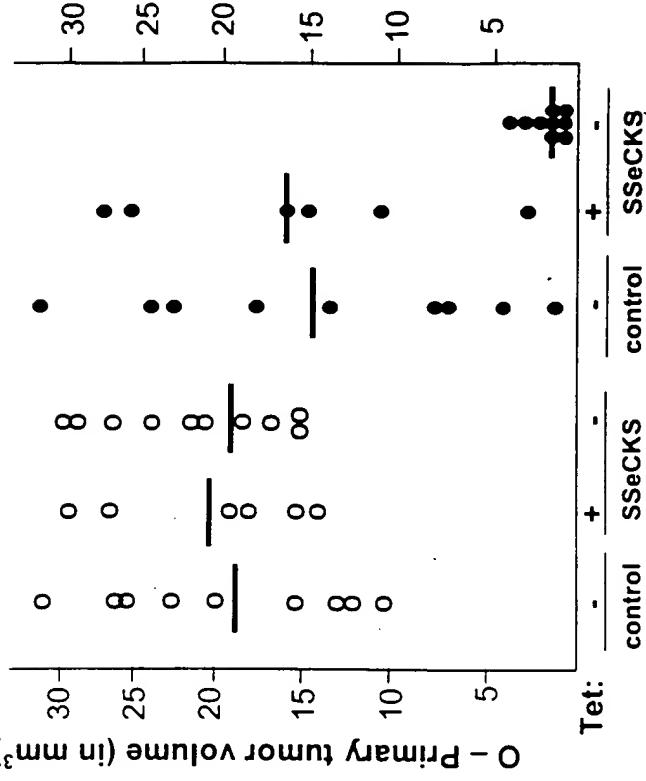


Figure 48 (78 of 90)

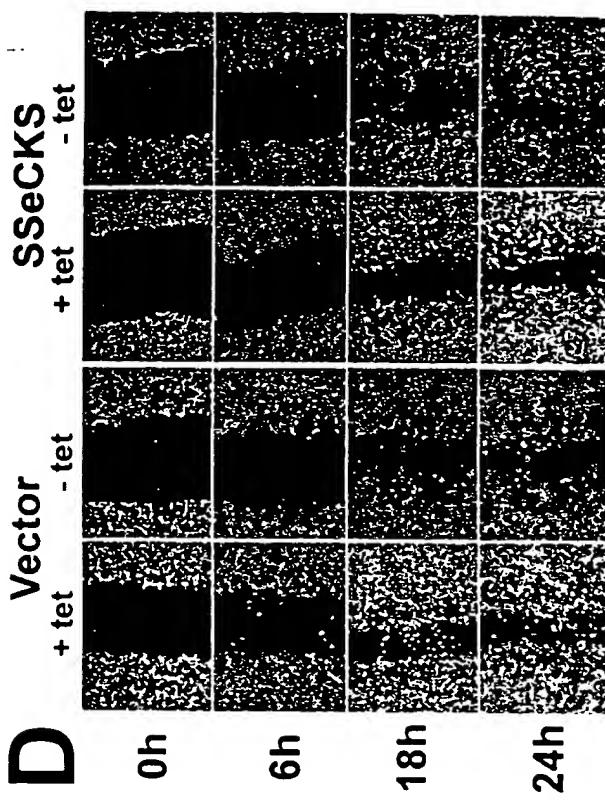
B



A



D



C v-2 2-6 2-6 v-2(T) 2-6(T)

tet: - + - + / - + / -



Figure 49

(79 of 90)

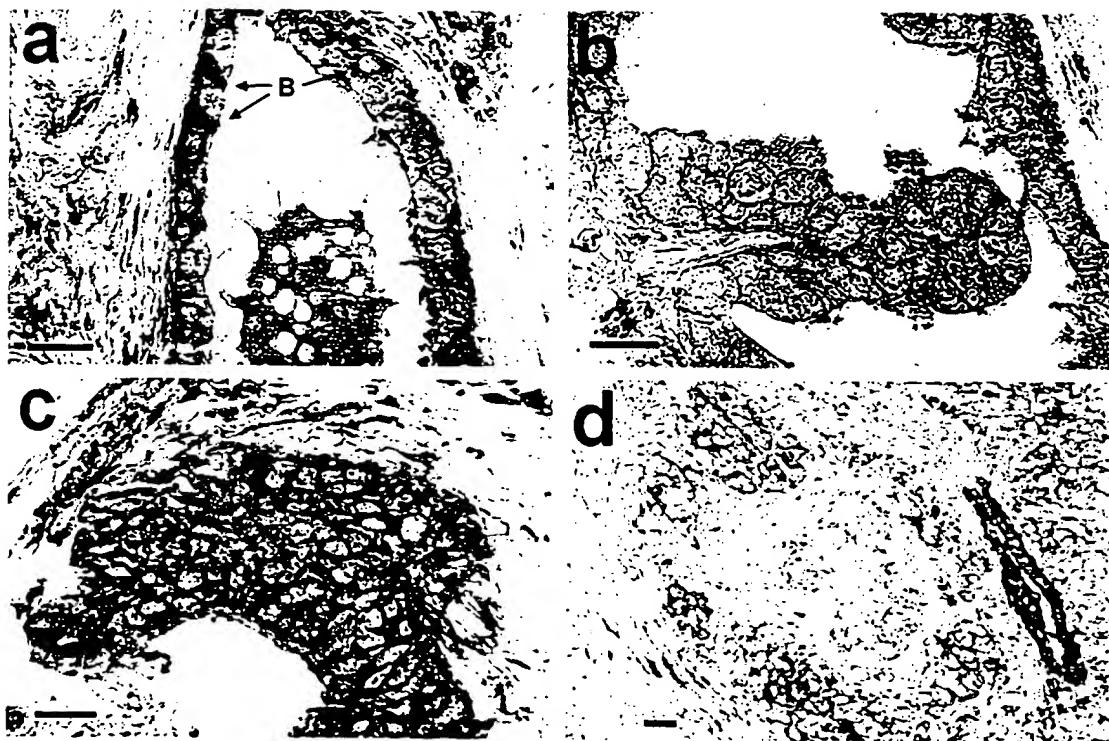


Figure 50

(80 of 90)

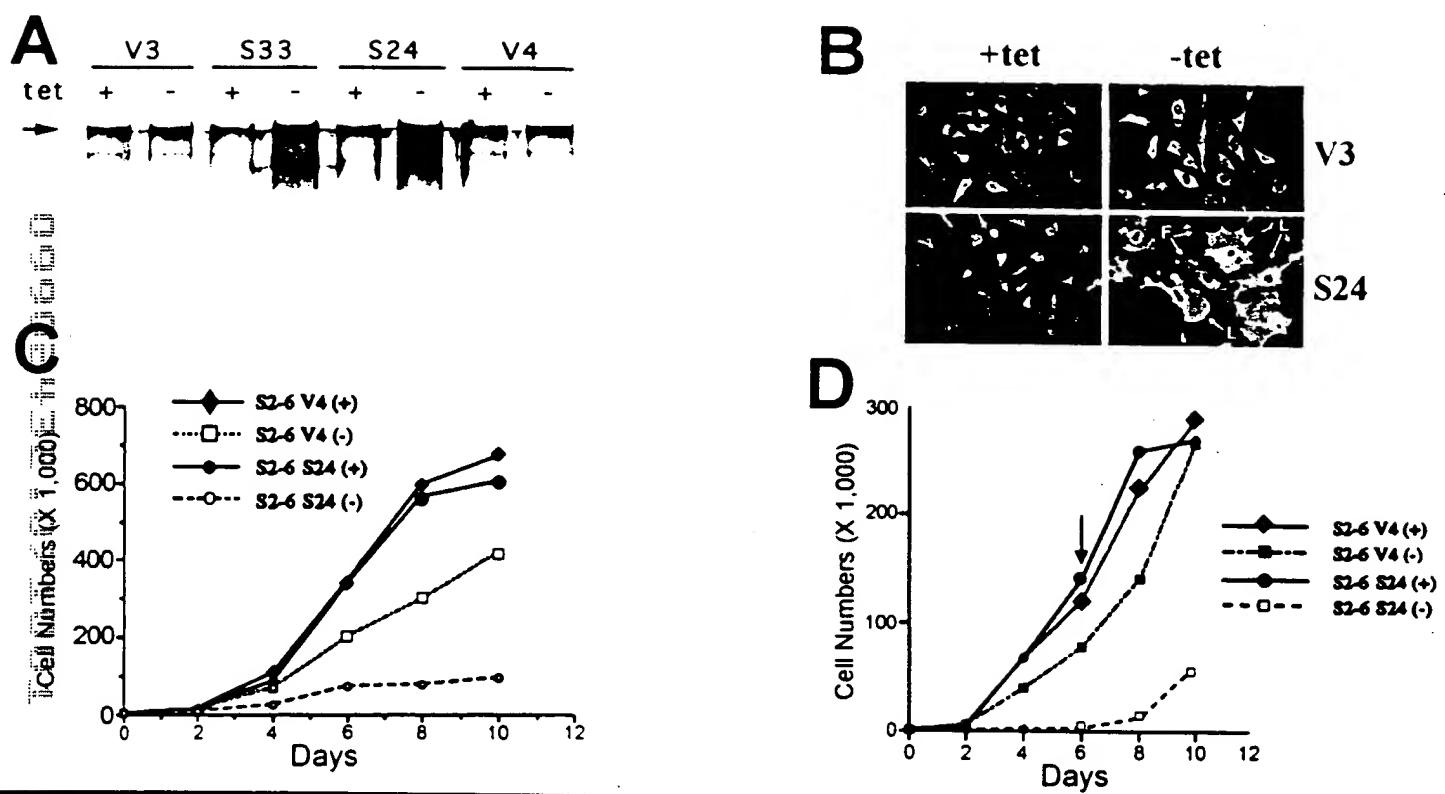


Figure 51

(81 of 90)

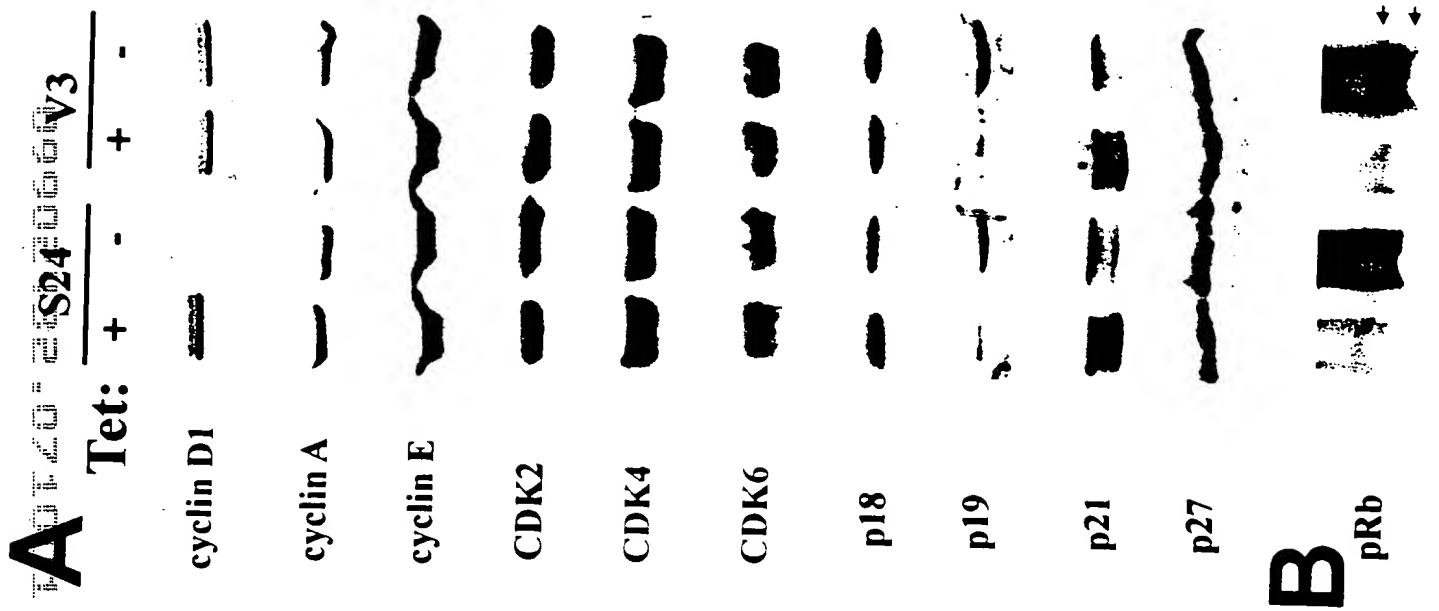


Figure 52

(82 of 90)

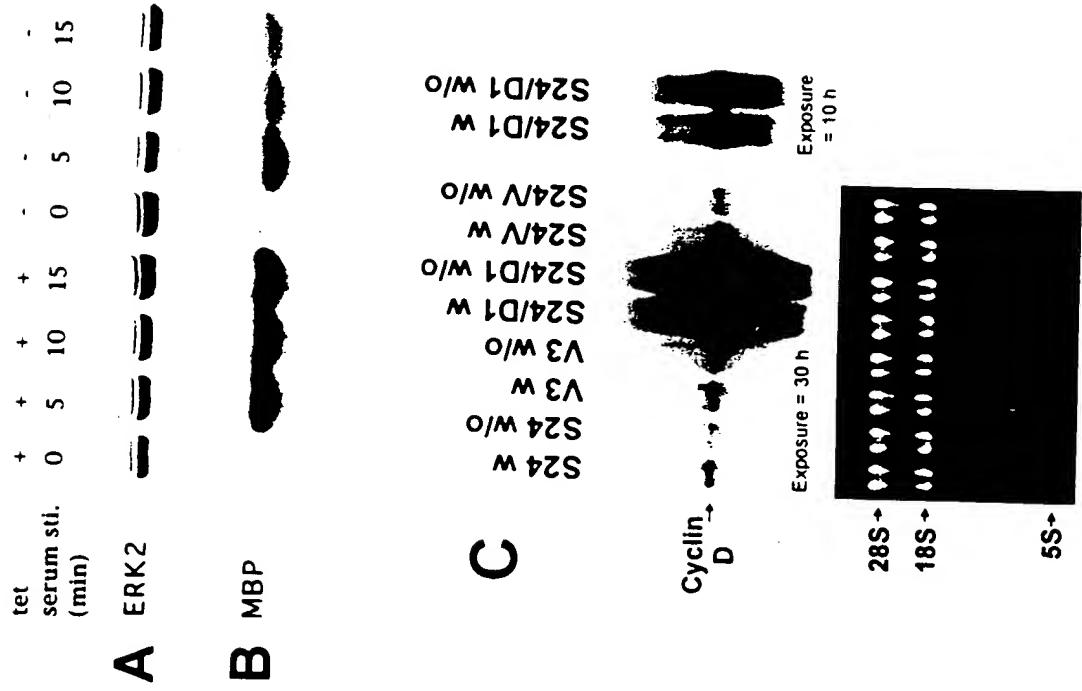


Figure 53

(83 of 90)

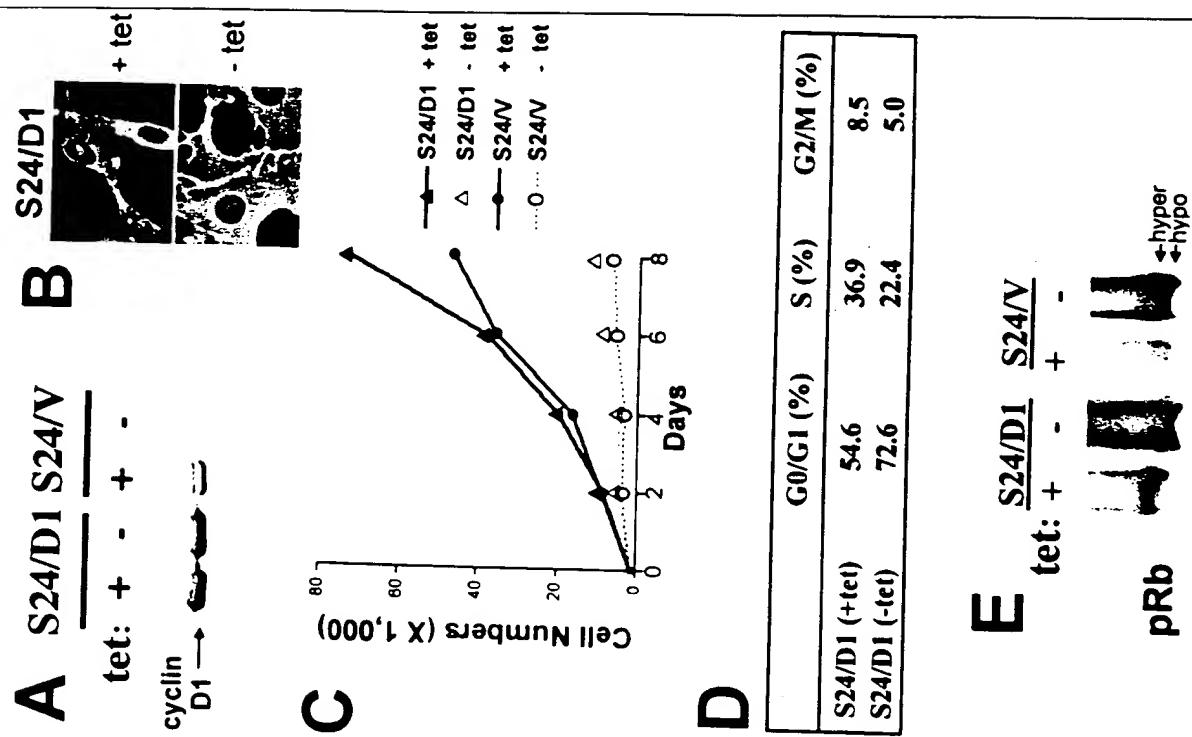


Figure 54

(84 of 90)

SSeCKS 468 SPEEKTLPKHPEGIVSEVEM LSSQERIK₄₉₆
Newt pRb || ||: |||||::| ||| ||::
780 SP. LKSPYKHPEGLLSPTKM - (27 a.a.) - LSSSERLR₈₃₄

Figure 55

85 of 90

A

S24/D1
(+tet)



S24/D1
(-tet)



V3/D1
(+tet)



V3/D1
(-tet)



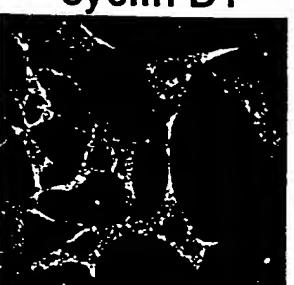
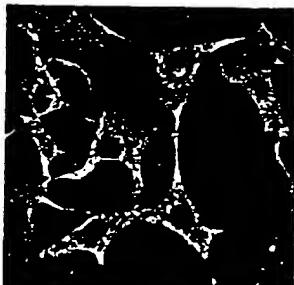
B

SSeCKS

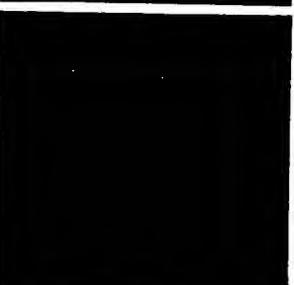
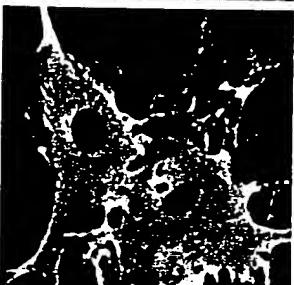
cyclin D1

SSeCKS +
cyclin D1

+ tet



- tet



C

- tet



Figure 56

(86 of 90)

S24/D1
+ tet



S24/D1
- tet



S24/D1
+ tet
+ PMA



S24/D1
- tet
+ PMA

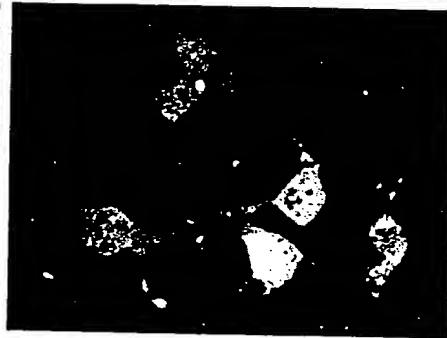


Figure 57

(87 of 90)

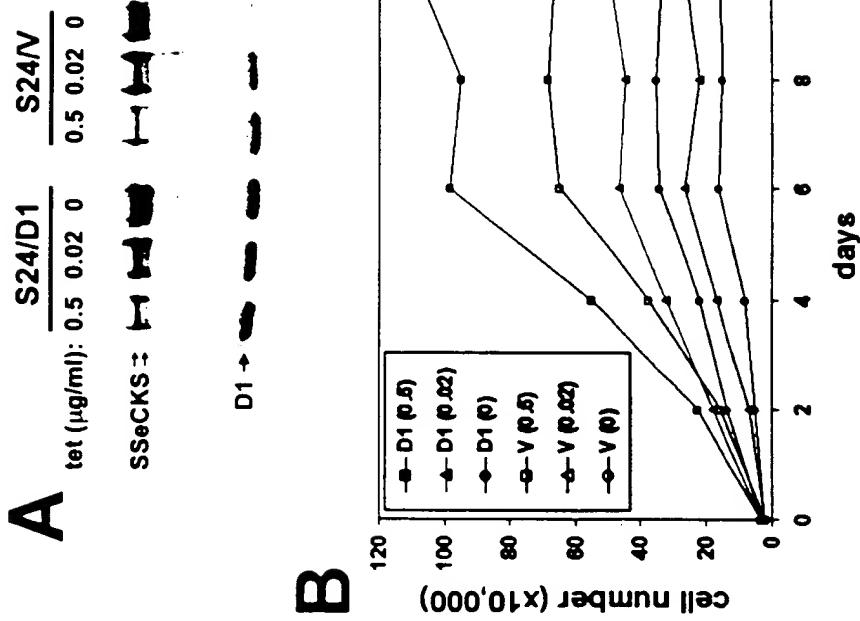


Figure 58

(88 of 90)

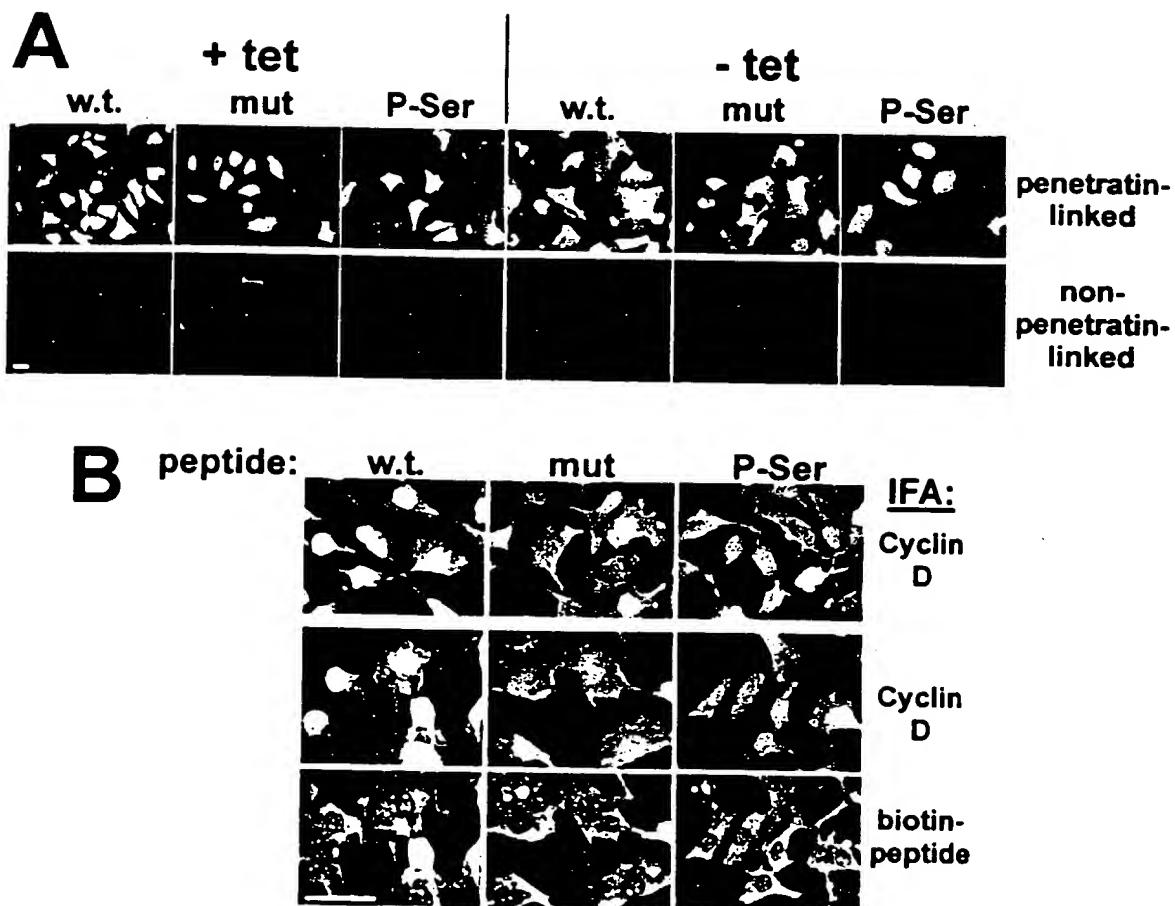
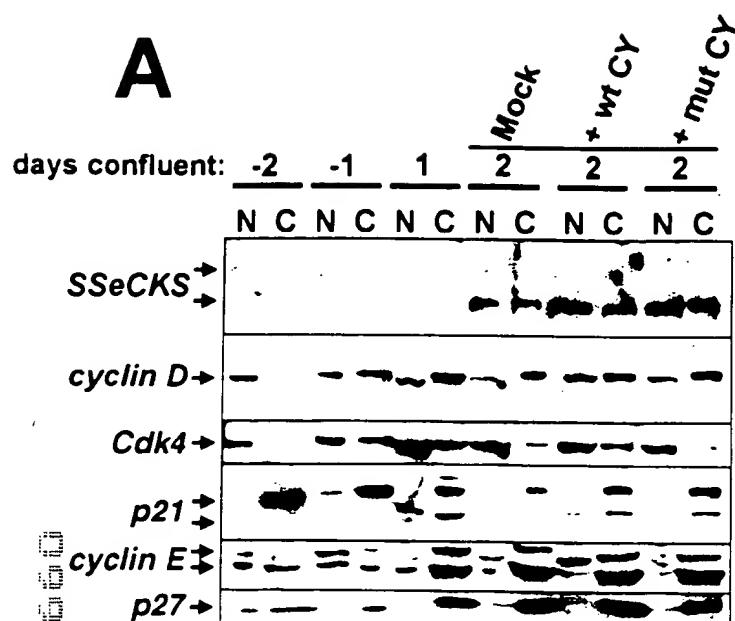


Figure 59

(89 of 90)

A



B

days confluent:

-2 -1 1 2 2 2

Mock
wt CY
mut CY

□ Nuclear
■ Cytoplasmic

SSeCKS

Cyclin D

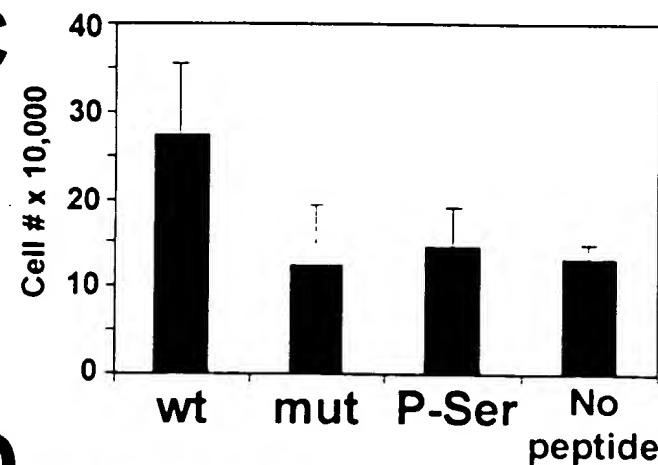
Cdk4

p21

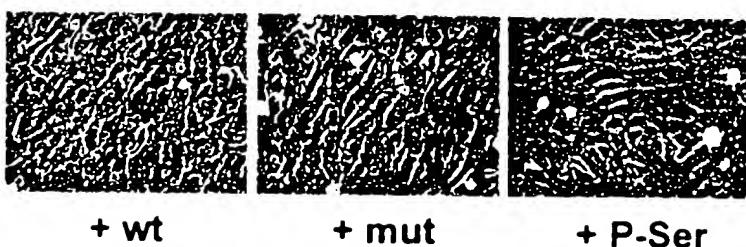
Cyclin E

p27

Relative protein levels



D



94A3

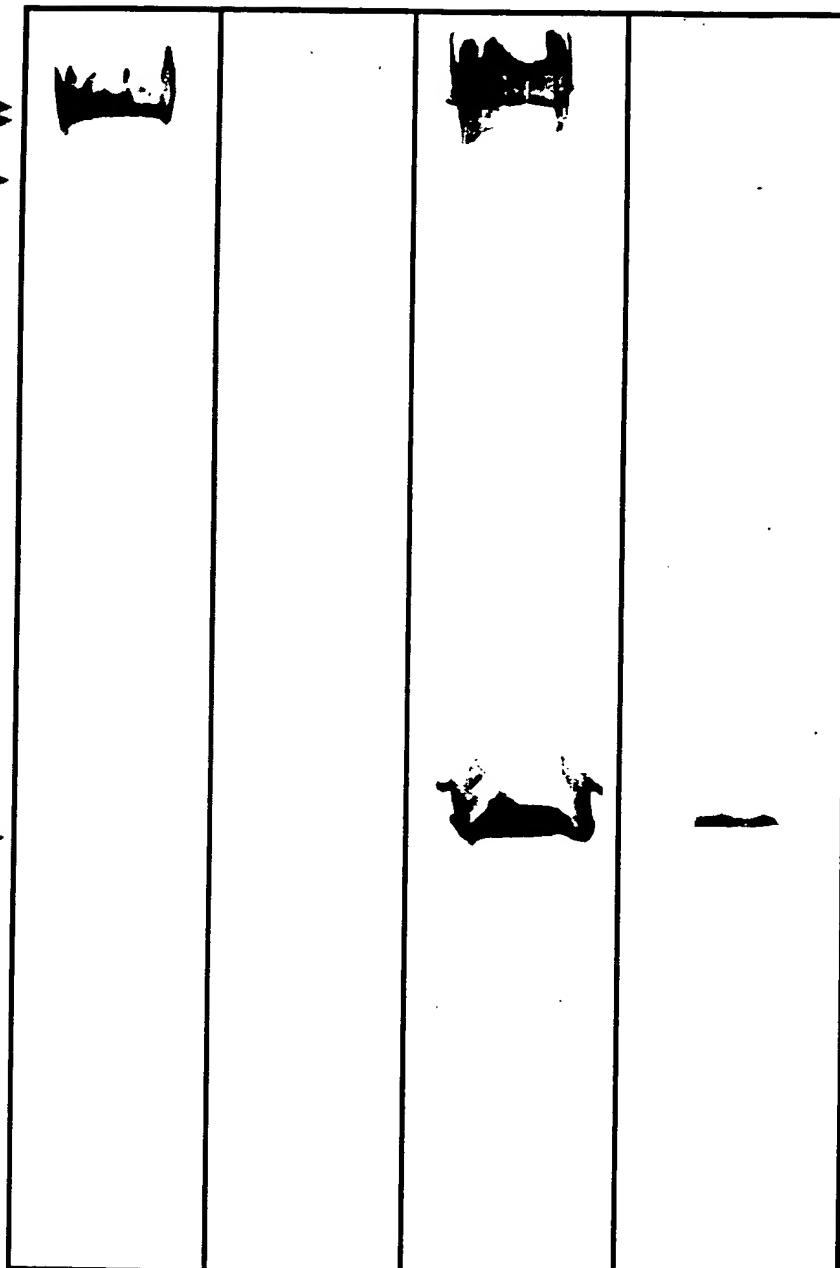
78H11

82B3

31A3

E

200kDa
100kDa
70kDa
55kDa
44kDa
30kDa
21kDa



200kDa

97.5

66

44

30

21